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(54) Title: APOPTOSIS MODULATORS THAT INTERACT WITH THE HUNTINGTON'S DISEASE GENE**(57) Abstract**

A family of proteins, including a specific human protein designated as HIP1, has been identified that interact differently with the gene product of a normal (16 CAG repeat) and an expanded (>44 CAG repeat) HD gene. Expression of the HIP1 protein was found to be enriched in the brain. Analysis of the sequence of the HIP1 protein indicated that it includes a death effector domain (DED), suggesting an apoptotic function. Thus, it appears that a normal function of Huntingtin may be to bind HIP1 and related apoptosis modulators, reducing its effectiveness in stimulating cell death. Since expanded huntingtin performs this function less well, there is an increase in HIP1-modulated cell death in individuals with an expanded repeat in the HD gene. This understanding of the likely role of huntingtin and HIP1 or related proteins (collectively "HIP-apoptosis modulating proteins") in the pathology of Huntington's disease offers several possibilities for therapy. First, because the function of huntingtin apparently depends at least in part on the ability to interact with HIP-apoptosis modulating proteins, added expression (e.g., via gene therapy) of normal (non-expanded) huntingtin or of the HIP-binding region of huntingtin should provide a therapeutic benefit. Other DED-interacting peptides could also be used to mask and reduce the interaction of HIP-apoptosis modulating proteins with the death signaling complex. Alternatively, a mutant form of HIP-protein from which the DED has been deleted might be introduced, for example using gene therapy techniques. Because HIP-apoptosis modulating proteins have been shown to self-associate, a protein with a deleted DED may compete with endogenous HIP-protein in the formation of these associations, thereby reducing the amount of apoptotically-active HIP-protein.

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APOPTOSIS MODULATORS THAT INTERACT WITH THE HUNTINGTON'S DISEASE GENE

BACKGROUND OF THE INVENTION

This application relates to a family of apoptosis modulators that interact with the Huntington's Disease gene product, and to methods and compositions relating thereto.

"Interacting proteins" are proteins which associate *in vivo* to form specific complexes.

5 Non-covalent bonds, including hydrogen bonds, hydrophobic interactions and other molecular associations form between the proteins when two protein surfaces are matched or have affinity for each other. This affinity or match is required for the recognition of the two proteins, and the formation of an interaction. Protein-protein interactions are involved in the assembly of enzyme subunits; in antigen-antibody reactions; in forming the supramolecular
10 structures of ribosomes, filaments, and viruses; in transport; and in the interaction of receptors on a cell with growth factors and hormones.

Huntington's disease is an adult onset disorder characterized by selective neuronal loss in discrete regions of the brain and spinal chord that lead to progressive movement disorder, personality change and intellectual decline. From onset, which generally occurs around age
15 40, the disease progresses with worsening symptoms, ending in death approximately 18 years after onset.

The biochemical cause of Huntington's disease is unclear. While the biochemical cause of Huntington's disease has remained elusive, a mutation in a gene within chromosome 4p16.3 subband has been identified and linked to the disease. This gene, referred to as the
20 Huntington's Disease or HD gene, contains two repeat regions, a CAG repeat region and a CCG repeat region. Testing of Huntington's disease patients has shown that the CAG region is highly polymorphic, and that the number of CAG repeat units in the CAG repeat region is a very reliable indicator of having inherited the gene for Huntington's disease. Thus, in control individuals and in most individuals suffering from neuropsychiatric disorders other than
25 Huntington's disease, the number of CAG repeats is between 9 and 35, while in individuals suffering from Huntington's disease the number of CAG repeats is expanded and is 36 or greater.

To date, no differences have been observed at either the total RNA, mRNA or protein levels between normal and HD-affected individuals. Thus, the function of the HD protein and its role in the pathogenesis of Huntington's Disease remain to be elucidated.

5 SUMMARY OF THE INVENTION

We have now identified a protein designated as HIP1, that interact differently with the gene product of a normal (16 CAG repeat) and an expanded (>44 CAG repeat) HD gene. The HIP1 protein originally isolated from a yeast two-hybrid screen is encoded by a 1.2 kb cDNA (Seq. ID. No. 1), devoid of stop codons, that is expressed as a 400 amino acid
10 polypeptide (Seq. ID. No. 2). Subsequent study has elucidated additional sequence for HIP1 such that a 1090 amino acid protein is now known. (Seq. ID No. 5). Expression of the HIP1 protein was found to be enriched in the brain.

Analysis of the sequence of the HIP1 protein indicated that it includes a death effector domain (DED), suggesting an apoptotic function. Thus, it appears that a normal function of
15 huntingtin may be to bind HIP1 and related apoptosis modulators, reducing its effectiveness in stimulating cell death. Since expanded huntingtin performs this function less well, there is an increase in HIP1-modulated cell death in individuals with an expanded repeat in the HD gene. Furthermore, additional members of the same family of proteins have been identified which also contain a DED. Thus, the present invention provides a new class of apoptotic
20 modulators which are referred to as HIP-apoptosis modulating proteins.

This understanding of the likely role of huntingtin and HIP1 or related proteins in the pathology of Huntington's Disease offers several possibilities for therapy. First, because the function of huntingtin apparently depends at least in part on the ability to interact with HIP-apoptosis modulating proteins, added expression (e.g., via gene therapy) of normal (non-
25 expanded) huntingtin or of the HIP-binding region of huntingtin should provide a therapeutic benefit. Other DED-interacting peptides could also be used to mask and reduce the interaction of HIP-apoptosis modulating proteins with the death signaling complex. Alternatively, a mutant form of HIP-protein from which the DED has been deleted might be introduced, for example using gene therapy techniques. Because HIP-apoptosis modulating
30 proteins have been shown to self-associate, a protein with a deleted DED may compete with

endogenous HIP-protein in the formation of these associations, thereby reducing the amount of apoptotically-active HIP-protein.

BRIEF DESCRIPTION OF THE DRAWING

5 Fig. 1 graphically depicts the amount of interaction between HIP1 and Huntingtin proteins with varying lengths of polyglutamine repeat;

 Fig. 2 compares the nucleic acid sequences of human and murine HIP1 and HIP1a;

 Fig. 3 compares the amino acid sequences of human and murine HIP1 and HIP1a;

10 Fig. 4 shows the sequences of various death effector domains in comparison to the DED of human and murine HIP1 and HIP1a;

 Fig. 5 shows the genomic organization of human HIP1;

 Fig. 6 compares the sequences of human HIP1 with ZK370.3 protein of *C. elegans*;

 Fig. 7 shows mouse EST's with homology to human HIP1 cDNA used to screen a mouse brain library;

15 Fig. 8 shows the affect of HIP1 on susceptibility of cells to stress; and

 Figs. 9A - 9C show the toxicity of HIP1 in the presence of huntingtin with different lengths of polyglutamine repeats.

DETAILED DESCRIPTION OF THE INVENTION

20 This application relates to a new family of proteins function as modulators of apoptosis. At least some of these proteins, notably the human protein designated HIP1, interact with the gene product of the Huntington's disease gene. Other proteins within the family possess at least 40% and preferably more than 50% nucleotide identity with HIP1 and include a death effector domain (DED) . Such proteins are referred to in the specification and claims
25 hereof as "HIP-apoptosis modulating proteins."

 The first HIP-apoptosis modulating protein identified was designated as HIP1. HIP1 was identified using the yeast two-hybrid system described in US Patent No. 5,283,173 which is incorporated herein by reference. Briefly, this system utilizes two chimeric genes or plasmids expressible in a yeast host. The yeast host is selected to contain a detectable marker
30 gene having a binding site for the DNA binding domain of a transcriptional activator. The

first chimeric gene or plasmid encodes a DNA-binding domain which recognizes the binding site of the selectable marker gene and a test protein or protein fragment. The second chimeric gene or plasmid encodes for a second test protein and a transcriptional activation domain.

The two chimeric genes or plasmids are introduced into the host cell and expressed, and the

5 cells are cultivated. Expression of the detectable marker gene only occurs when the gene product of the first chimeric gene or plasmid binds to the DNA binding domain of the detectable marker gene, and a transcriptional activation domain is brought into sufficient proximity to the DNA-binding domain, an occurrence which is facilitated by protein-protein interactions between the first and second test proteins. By selecting for cells expressing the
10 detectable marker gene, those cells which contain chimeric genes or plasmids for interacting proteins can be identified, and the gene can be recovered and identified.

In testing for Huntington Interacting Proteins, several different plasmids were prepared containing portions of the human HD gene. The first four, identified as 16pGBT9, 44pGBT9, 80pGBT9 and 128pGBT9, were GAL4 DNA binding domain-HD in-frame
15 fusions containing nucleotides 314 to 1955 (amino acids 1-540) of the published HD cDNA sequences cloned into the vector pGBT9 (Clontech). These plasmids contain a CAG repeat region of 16, 44, 80 and 128 glutamine-encoding repeats, respectively. A clone (DMK BamHlpGBT9) was made by fusing a cDNA encoding the first 544 amino acids of the myotonic dystrophy gene (a gift from R. Korneluk) in-frame with the GAL4-DNA BD of
20 pGBT9 and was used as a negative control.

These plasmids have been used to identify and characterize HIP1, as well as two additional HD-interacting proteins, HIP2 and HIP3, which have not yet been tested for function as apoptosis modulators. These plasmids can be further used for the identification of additional interacting proteins which do act as apoptosis modulators, and for tests to refine
25 the region on the protein in which the interaction occurs. Thus, one aspect of the invention is these four plasmids, and the use of these plasmids in identifying HD-interacting proteins. Furthermore, it will be appreciated that the GAL4 DNA-binding and activating domains are not the only domains which can be used in the yeast two-hybrid assay. Thus, in a broader sense, the invention encompasses any chimeric genes or plasmids containing nucleotides 314
30 to 1955 of the HD gene together with an activating or DNA-binding domain suitable for use

in the yeast one, two- or three-hybrid assay for proteins critical in either binding to the HD protein or responsible for regulated expression of the HD gene.

After introducing the plasmids into Y190 yeast host cells, transforming the host cells with an adult human brain Matchmaker™ (Clontech) cDNA library coupled with a GAL4 activating domain, and selecting for the expression of two detectable marker genes to identify clones containing genes for interacting proteins, the activating domain plasmids were recovered and analyzed. As a result of this analysis, three different cDNA fragments were identified as encoding for HD-interacting proteins and designated as HIP1, HIP2 and HIP3. The nucleic acid sequence of HIP1, as originally recovered in the yeast two-hybrid assay, is given in Seq. ID. No 1. The polypeptide which it encodes is given by Seq. ID No. 2. Further investigation of the HIP1 cDNA resulted in the characterization of a longer region of cDNA totaling 4795 bases and a corresponding protein, the sequences of which are given by Seq ID Nos. 3 and 4, respectively. A further portion of the HIP1 protein was characterized, extending the length to the complete protein sequence of 1090 amino acids (Seq. ID No. 5)

The cDNA molecules encoding HIP-apoptosis modulating proteins, particularly those encoding portions of HIP1, can be explored using oligonucleotide probes for example for amplification and sequencing. In addition, oligonucleotide probes complementary to the cDNA can be used as diagnostic probes to localize and quantify the presence of HIP1 DNA. Probes of this type with a one or two base mismatch can also be used in site-directed mutagenesis to introduce variations into the HIP1 sequence which may increase or decrease the apoptotic activity. Preferred targets for such mutations would be the death effector domains. Thus, a further aspect of the present invention is an oligonucleotide probe, preferably having a length of from 15-40 bases which specifically and selectively hybridizes with the cDNA given by Seq. ID No. 1 or 3 or a sequence complementary thereto. As used herein, the phrase "specifically and selectively hybridizes with" the cDNA refers to primers which will hybridize with the cDNA under stringent hybridization conditions.

Probes of this type can also be used for diagnostic purposes to characterize risk of Huntington's Disease like symptoms arising in individuals where the symptoms are present in the family history but are not associated with an expansion of the CAG repeat. Such symptoms may arise from a mutation in HIP1 or other HIP-apoptosis modulating protein

which alters the interaction of this protein with huntingtin, thereby increasing the apoptotic activity of the protein even in the presence of a normal (non-expanded) huntingtin molecule. An appropriate probe for this purpose would one which hybridizes with or adjacent to the huntingtin binding region of the HIP-apoptosis modulating protein. In HIP1, this lies within
5 amino acids 129-514.

DNA sequencing of the HIP1 cDNA initially isolated from the yeast two-hybrid screen (Seq. ID No. 1) revealed a 1.2 kb cDNA that shows no significant degree of nucleic acid identity with any stretch of DNA using the blastn program at ncbi (blast@ncbi.nlm.nih.gov). When the larger HIP1 cDNA sequence (SEQ ID NO. 3) was
10 translated into a polypeptide, the HIP1 cDNA coding (nucleotides 328-3069) is observed to be devoid of stop codons, and to produce a 914 amino acid polypeptide. A polypeptide identity search revealed an identity match over the entire length of the protein (46% conservation) with that of a hypothetical protein from *C. elegans* (ZK370.3 protein; *C. elegans* cosmid ZK370). This *C. elegans* protein shares identity with the mouse talin gene,
15 which encodes a 217 kDa protein implicated with maintaining integrity of the cytoskeleton. It also shares identity with the SLA2/MOP2/ END4 gene from *Saccharomyces cerevisiae*, which is known to code for an essential cytoskeletal associated gene required for the accumulation and or maintenance of plasma membrane H⁺-ATPase on the cell surface. When pairwise comparisons are performed between HIP1 and the *C. elegans* ZK370.3 protein
20 (Genpept accession number celzk370.3), it shows 26% complete identity and an overall 46% level of conservation. Comparative analysis between HIP1 and SLA2/MOP2/ END4 (EMBL accession number Z22811) demonstrate similar conservation (20% identity, 40% conservation).

Further exploration revealed several important facts about HIP1 that implicate it in a
25 significantly in the pathogenesis of Huntington's Disease. First, as shown in Fig. 1, it was found that the native interaction between HD protein and HIP1 is influenced by the number of CAG repeats. Second, it was found that expression of the HIP1 protein is enriched in the brain. The highest amounts of expression are in the cortex, with lower levels being seen in the cerebellum, caudate and putamen.

It has also been observed that huntingtin proteins with expanded polyglutamine tracts can aggregate into large, irregularly shaped deposits in HD brains, transgenic mice and *in vitro* cell culture. We have shown that in HEK (human embryonic kidney) 293T cells, the aggregation of full-length and smaller huntingtin fragments occurs after the cells have been exposed to a period of apoptotic stress. Martindale, et al., *Nature Genetics* 18: 150-154 (1998). In order to assess the consequence of HIP1 expression in cultured cells, we used huntingtin aggregation as one marker of viability. What we found was that cells cotransfected with huntingtin (128 CAG repeats) and HIP1 contained aggregates comparable to those observed following application of apoptotic stress with sub-lethal doses of tamoxifen in 14% of the cells, and that these cells were the ones in which both genes had been introduced as reflected by a double marker experiment. Transfection of a gene encoding a fusion protein of 128 repeat huntingtin and the DED domain from HIP1 ligated in the sense orientation resulted in aggregate formation in 30 to 50% of the cells.

The implications of the apoptotic activity of HIP1 are two-fold. First, the fact that this activity is apparently differentially modulated by interaction with huntingtin having normal and expanded repeats implicates HIP1 in the apoptotic neuronal death which is observed in Huntington's disease and makes HIP1 a logical target for therapy. A second implication of the apoptotic activity of HIP1 is the potential for use of HIP1 as a therapeutic agent to introduce apoptosis in cancer cells.

Therapeutic targeting of HIP1 or other HIP-apoptosis modulating proteins might take any of several forms, but will in general be a treatment involving administration of a composition that reduces the apoptotic activity of the HIP-apoptosis modulating protein. As used in the specification and claims hereof, the term "administration" includes direct administration of a composition active to reduce apoptotic activity as well as indirect administration which might include administration of pro-drugs or nucleic acids that encode the desired therapeutic composition.

One class of composition which can be used in the therapeutic methods of the invention are those compositions which interfere with the activity of HIP-apoptosis modulating proteins by binding to the proteins and mask and reduce the interaction of HIP-apoptosis modulating proteins with the death signaling complex. Within this class of

compositions are normal (non-expanded) huntingtin, administered, for example, via increased expression of exogenous HD genes; the HIP-binding region of huntingtin, administered via gene therapy techniques; and other DED-interacting peptides. Other DED-interacting peptides which might be used in a therapeutic method of this type include FADD (Beldin et al., *Cell* 85: 803-815 (1996)) and caspase 8 (Muzio et al., *Cell* 85: 817-827 (1996)).

An alternative form of therapy involves the use of a mutant form of HIP1 or other HIP-apoptosis modulating protein from which the DED has been deleted. DED-containing proteins, including HIP1 are self-associating, and this self-association has been shown to be important for activity. (Muzio et al., *Cell* 85: 817-827 (1996)). Thus, a protein with a deleted DED may compete with endogenous HIP-protein in the formation of these associations, thereby reducing the amount of apoptotically-active HIP-protein.

In addition to HIP1, we have identified a further human protein, HIP1a, from a human frontal cortex cDNA library. HIP1a is a family member of HIP1, and thus a HIP-apoptosis modulator in accordance with the invention. A partial sequence of HIP1a (the 5' portion of HIP1a remains to be characterized) is given by SEQ ID Nos. 6 and 7. The isolated and characterized portion of HIP1a shows 53% nucleotide identity and 58% amino acid conservation with HIP1 (Table 1, Figs. 2 and 3).

We have also isolated 2 mouse proteins mHIP1 and mHIP1a (SEQ. ID Nos. 8-11) which appear to be the murine homologues of human HIP1 and HIP1a. As in the case of human HIP1a, the 5' portion of mHIP1 remains to be isolated. At present, mHIP1 shows 85% nucleotide identity and 90% amino acid conservation with huHIP1 (Table 1, Figs. 2 and 3). mHIP1a shows 60% nucleotide identity and 61% amino acid conservation with huHIP1 (Table 1, Figs. 2 and 3). mHIP1a shows stronger homology to huHIP1a; it shows 87% nucleotide identity and 91% amino acid conservation with huHIP1a (Table 1, Figs. 2 and 3). Taken together these findings indicate that mHIP1 is the murine homologue of huHIP1 whereas mHIP1a is most likely the murine homologue of huHIP1a. As mentioned previously, HIP1 shows sequence similarity to Sla2p in *S. cerevisiae* and the hypothetical protein ZK370.3 in *C. elegans*. Similarly, huHIP1a, mHIP1, and mHIP1a show sequence similar to Sla2p and ZK370.3 (Table 2). The carboxy-terminal regions of huHIP1a, mHIP1, and mHIP1a all show considerable homology to the mammalian membrane

cytoskeletal-associated protein, talin. This suggests that these 3 proteins may also play a role in the regulation of membrane events through interactions with the underlying cytoskeleton.

HIP1 contains a death effector domain (DED), a domain which is also present in a number of proteins involved in the apoptotic pathway (Fig. 4). This suggests that HIP1 may
5 act as a modulator of the apoptosis pathway. The DED in huHIP1 is present between amino acid positions 287 and 368. Similarly, HIP1a, mHIP1, and mHIP1a also contain a DED. In huHIP1a the DED is present at amino acids 1-78 of the recovered fragment. In mHIP1 and mHIP1a, the DED are present at amino acids 128- 210 and 388-470, respectively. The DED present in huHIP1a, mHIP1 and mHIP1a all show significant percentage amino acid
10 conservation to the DED present in huHIP1 (Table 3).

Increasing expression of normal (non-expanded) huntingtin or the HIP-apoptotic modulator-binding portion thereof, a modified HIP-apoptotic modulator in which the DED has been deleted or of a DED-interacting protein or peptide can be accomplished using gene therapy approaches. In general, this will involve introduction of DNA encoding the
15 appropriate protein or peptide in an expressable vector into the brain cells. Expression of HIP-apoptosis modulating proteins may also be useful in treatment of cancer in which case application to other cell types would be desired, and cells expressing HIP-apoptosis modulating proteins may be used for screening of therapeutic compounds. Thus, in a more general sense, expression vectors are defined herein as DNA sequences that are required for
20 the transcription of cloned copies of genes and the translation of their mRNAs in an appropriate cell type. Specifically designed vectors allow the shuttling of DNA between hosts such as bacteria-yeast or bacteria-animal cells. An appropriately constructed expression vector may contain: an origin of replication for autonomous replication in host cells, selectable markers, a limited number of useful restriction enzyme sites, a potential for high
25 copy number, and active promoters. A promoter is defined as a DNA sequence that directs RNA polymerase to bind to DNA and initiate RNA synthesis. A strong promoter is one which causes mRNAs to be initiated at high frequency. Expression vectors may include, but are not limited to, cloning vectors, modified cloning vectors, specifically designed plasmids or viruses.

30 A variety of mammalian expression vectors may be used to express recombinant

HIP-apoptosis modulating proteins or fragments thereof in mammalian cells. Commercially available mammalian expression vectors which may be suitable for recombinant HIP-apoptosis modulating protein expression, include but are not limited to, pMC1neo (Stratagene), pXT1 (Stratagene), pSG5 (Stratagene), EBO-pSV2-neo (ATCC 37593) 5 pBPV-1(8-2) (ATCC 37110), pdBPV-MMTneo(342-12) (ATCC 37224), pRSVgpt (ATCC 37199), pRSVneo (ATCC 37198), pSV2-dhfr (ATCC 37146), pUCTag (ATCC 37460), and IZD35 (ATCC 37565). Other vectors which have been shown to be suitable expression systems in mammalian cells include the herpes simplex viral based vectors: pHSV1 (Geller et al. Proc. Natl. Acad. Sci. 87:8950-8954 (1990)); recombinant retroviral vectors: MFG 10 (Jaffee et al. Cancer Res. 53:2221-2226 (1993)); Moloney-based retroviral vectors: LN, LNSX, LNCX, LXSX (Miller and Rosman Biotechniques 7:980-989 (1989)); vaccinia viral vector: MVA (Sutter and Moss Proc. Natl. Acad. Sci. 89:10847-10851 (1992)); recombinant adenovirus vectors : pJM17 (Ali et al Gene Therapy 1:367-384 (1994)), (Berkner K. L. Biotechniques 6:616-624 1988); second generation adenovirus vector: DE1/DE4 adenoviral 15 vectors (Wang and Finer Nature Medicine 2:714-716 (1996)); and Adeno-associated viral vectors: AAV/Neo (Muro-Cacho et al. J. Immunotherapy 11:231-237 (1992)).

The expression vector may be introduced into host cells via any one of a number of techniques including but not limited to transformation, transfection, infection, protoplast fusion, and electroporation. The expression vector-containing cells are clonally propagated 20 and individually analyzed to determine whether they produce the desired protein. Delivery of retroviral vectors to brain and nervous system tissue has been described in US Patents Nos. 4,866,042, 5,082,670 and 5,529,774, which are incorporated herein by references. These patents disclose the use of cerebral grafts or implants as one mechanism for introducing vectors bearing therapeutic gene sequences into the brain, as well as an approach in which the 25 vectors are transmitted across the blood brain barrier.

To further illustrate the methods of making the materials which are the subject of this invention, and the testing which has established their utility, the following non-limiting experimental procedures are provided.

EXAMPLE 1

IDENTIFICATION OF INTERACTING PROTEINS

GAL4-HD cDNA constructs

An HD cDNA construct (44pGBT9), with 44 CAG repeats was generated encompassing amino acids 1 - 540 of the published HD cDNA . This cDNA fragment was fused in frame to the GAL4 DNA-binding domain (BD) of the yeast two-hybrid vector pGBT9 (Clontech). Other HD cDNA constructs, 16pGBT9, 80pGBT9 and 128pGBT9 were constructed, identical to 44pGBT9 but included only 16, 80 or 128 CAG repeats, respectively.

Another clone (DMKDBamHlpGBT9) containing the first 544 amino acids of the myotonic dystrophy gene (a gift from R. Korneluk) was fused in-frame with the GAL4-DNA BD of pGBT9 and was used as a negative control. Plasmids expressing the GAL4-BDRAD7 (D. Gietz, unpublished) and SIR3 were used as a positive control for the β -galactosidase filter assay.

The clones IT15-23Q, IT15-44Q and HAP1 were generous gifts from Dr. C. Ross. These clones represent a previously isolated huntingtin interacting protein that has a higher affinity for the expanded form of the HD protein.

Yeast strains, transformations and β -galactosidase assays

The yeast strain Y190 (MATa leu2-3,112, ura3-52, trp1-901, his3- Δ 200, ade2-101, gal4 Δ gal80 Δ , URA3::GAL-lacZ, LYS2::GAL-HIS3,cyc') was used for all transformations and assays. Yeast transformations were performed using a modified lithium acetate transformation protocol and grown at 30 C using appropriate synthetic complete (SC) dropout media.

The β -galactosidase chromogenic filter assays were performed by transferring the yeast colonies onto Whatman filters. The yeast cells were lysed by submerging the filters in liquid nitrogen for 15-20 seconds. Filters were allowed to dry at room temperature for at least five minutes and placed onto filter paper presoaked in Z-buffer (100 mM sodium phosphate (pH7.0) 10 mM KCl, 1 mM MgSO₄) supplemented with 50 mM

2-mercaptoethanol and 0.07 mg/ml 5-bromo-4-chloro-3-indolyl β -D-galactoside (X-gal). Filters were placed at 37 C for up to 8 hours.

Yeast two-hybrid screening for huntingtin interacting protein (HIP)

5 cDNAs from an human adult brain Matchmaker™ cDNA library (Clontech) was transformed into the yeast strain Y190 already harboring the 44pGBT9 construct. The transformants were plated onto one hundred 150 mm x 15 mm circular culture dishes containing SC media deficient in Trp, Leu and His. The herbicide 3-amino-triazole (3-AT) (25mM) was utilized to limit the number of false His⁺ positives (31). The yeast
10 transformants were placed at 30 C for 5 days and β -galactosidase filter assays were performed on all colonies found after this time, as described above, to identify β -galactosidase⁺ clones. Primary His⁺/ β -galactosidase⁺ clones were then orderly patched onto a grid on SC -Trp/-Leu/-His (25 mM 3AT) plates and assayed again for His⁺ growth and the ability to turn blue with a filter assay. Secondary positives were identified for further analysis. Proteins
15 encoded by positive cDNAs were designated as HIPs (Huntingtin Interactive Proteins). Approximately 4.0×10^7 Trp/Leu auxotrophic transformants were screened and of 14 clones isolated 12 represented the same cDNA (HIP1), and the other 2 cDNAs, HIP2 and HIP3 were each represented only once.

The HIP cDNA plasmids were isolated by growing the His⁺/ β -galactosidase⁺ colony
20 in SC -Leu media overnight, lysing the cells with acid-washed glass beads and electroporating the bacterial strain, KC8 (leuB auxotrophic) with the yeast lysate. The KC8 ampicillin resistant colonies were replica plated onto M9 (-Leu) plates. The plasmid DNA from M9⁺ colonies was transformed into DH5-a for further manipulation.

25

EXAMPLE 2

CONFIRMATION OF INTERACTIONS

The HIP1-GAL4-AD cDNA activated both the lac-Z and His reporter genes in the yeast strain Y190 only when co-transformed with the GAL4-BD-HD construct, but not the negative controls (Fig. 1) of the vector alone or a random fusion protein of the myotonin
30 kinase gene. In order to assess the influence of the polyglutamine tract on the interaction

between HIP1 and HD, semi-quantitative β -galactosidase assays were performed. GAL4-BD-HD fusion proteins with 16, 44, 80 and 128 glutamine repeats were assayed for their strength of interaction with the GAL4-AD-HIP1 fusion protein.

Liquid β -galactosidase assays were performed by inoculating a single yeast colony into appropriate synthetic complete (SC) dropout media and grown to OD600 0.6-1.5. Five millilitres of overnight culture was pelleted and washed once with 1 ml of Z-Buffer, then resuspended in 100 ml Z-Buffer supplemented with 38 mM 2-mercaptoethanol, and 0.05% SDS. Acid washed glass beads (~100 ml) were added to each sample and vortexed for four minutes, by repeatedly alternating a 30 seconds vortex, with 30 seconds on ice. Each sample was pelleted and 10 ml of lysate was added to 500 ml of lysis buffer. The samples were incubated in a 30 C waterbath for 30 seconds and then 100 ml of a 4 mg/ml o-nitrophenyl b-D galactopyranoside (ONPG) solution was added to each tube. The reaction was allowed to continue for 20 minutes at 30 C and stopped by the addition of 500 ml of 1 M Na₂CO₃ and placing the samples on ice. Subsequently, OD420 was taken in order to calculate the β -galactosidase activity with the equation $1000 \times \text{OD420} / (t \times V \times \text{OD600})$ where t is the elapsed time (minutes) and V is the amount of lysate used.

The specificity of the HIP1-HD interaction can be observed using the chromogenic filter assay. Only yeast cells harboring HIP1 and HD activate both the HIS and lacZ reporter genes in the Y190 yeast host. The cells that contain the HIP1 with HD constructs with 80 or 128 CAG repeats turn blue approximately 45 minutes after the cells with the smaller sized repeats (16 or 44).

No difference in the β -galactosidase activity was observed between the 16 and 44 repeats or between the 80 and 128 repeats. However, a significant difference ($p < 0.05$) in activity is seen between the smaller repeats (16 and 44) and the larger repeats (80 and 128). (Figure 1)

EXAMPLE 3

DNA SEQUENCING, cDNA ISOLATION AND 5' RACE

Oligonucleotide primers were synthesized on an ABI PCR-mate oligo-synthesizer. DNA sequencing was performed using an ABI 373 fluorescent automated DNA sequencer.

The HIP cDNAs were confirmed to be in-frame with the GAL4-AD by sequencing across the AD-HIP1 cloning junction using an AD oligonucleotide (5'GAA GAT ACC CCA CCA AAC3'). (Seq. ID No. 12)

5 Subsequently, primer walking was used to determine the remaining sequences. A human frontal cortex >4.0 kb cDNA library (a gift from S. Montal) was screened to isolate the full length HIP1 gene. Fifty nanograms of a 558 base pair Eco RI fragment from the original HIP1 cDNA was radioactively labeled with [α^{32} P]-dCTP using nick-translation and the probe allowed to hybridized to filters containing >105 pfu/ml of the cDNA library overnight at 65°C in Church buffer (see Northern blot protocol). The filters were washed at 10 65°C for 10 minutes with 1 X SSPE, 15 minutes at 65 C with 1 X SSPE and 0.1% SDS, then for thirty minutes and fifteen minutes with 1 X SSPE and 0.1% SDS. The filters were exposed to X-ray film (Kodak, XAR5) overnight at -70 C. Primary positives were isolated and replated and subsequent secondary positives were hybridized and washed as for the primary screen. The resulting positive phage were converted into plasmid DNA by 15 conventional methods (Stratagene) and the cDNA isolated and sequenced.

In order to obtain the most 5' sequence of the HIP1 gene, a Rapid Amplification of cDNA Ends (RACE) protocol was performed according to the manufacturers recommendations (BRL). First strand cDNA was synthesized using the oligo HIP1-242R (5' GCT TGA CAG TGT AGT CAT AAA GGT GGC TGC AGT CC 3'). (Seq. ID No. 13) 20 After dCTP tailing the cDNA with terminal deoxy transferase, two rounds of 35 cycles (94°C 1 minute; 53°C 1 minute; 72°C 2 minutes) of PCR using HIP1-R2 (5' GGA CAT GTC CAG GGA GTT GAA TAC 3') (Seq. ID No. 14) and an anchor primer (5' (CUA)4 GGC CAC GCG TCG ACT AGT ACG GGI IGG GII GGG IIG3') (BRL ,Seq. ID No. 15)) were performed. The subsequent 650 base pair PCR product was cloned using the TA 25 cloning system (Invitrogen) and sequenced using T3 and T7 primers. Sequences ID Nos. 1 and 3 show the sequence of the HIP1 cDNAs obtained.

EXAMPLE 4

DNA AND AMINO ACID ANALYSES

Overlapping DNA sequence was assembled using the program MacVector and sent via email or Netscape to the BLAST server at NIH (<http://www.ncbi.nlm.nih.gov>) to search for sequence similarities with known DNA (blastn) or protein (tblastn) sequences. Amino acid alignments were performed with the program Clustalw.

EXAMPLE 5

FISH DETECTION SYSTEM AND IMAGE ANALYSIS

The HIP1 cDNA isolated from the two-hybrid screen was mapped by fluorescent in situ hybridization (FISH) to normal human lymphocyte chromosomes counterstained with propidium iodide and DAPI. Biotinylated probe was detected with avidin-fluorescein isothiocyanate (FITC). Images of metaphase preparations were captured by a thermoelectrically cooled charge coupled camera (Photometrics). Separate images of DAPI banded chromosomes and FITC targeted chromosomes were obtained. Hybridization signals were acquired and merged using image analysis software and pseudo colored blue (DAPI) and yellow (FITC) as described and overlaid electronically. This study showed that HIP1 maps to a single genomic locus at 7q11.2.

EXAMPLE 6

NORTHERN BLOT ANALYSIS

RNA was isolated using the single step method of homogenization in guanidinium isothiocyanate and fractionated on a 1.0% agarose gel containing 0.6 M formaldehyde. The RNA was transferred to a hybond N -membrane (Amersham) and crosslinked with ultraviolet radiation.

Hybridization of the Northern blot with b-actin as an internal control probe provided confirmation that the RNA was intact and had transferred. The 1.2 kb HIP1 cDNA was labeled using nick translation and incorporation of $\alpha^{32}\text{P}$ -dCTP. Hybridization of the original 1.2 kb HIP1 cDNA was carried out in Church buffer (0.5 M sodium phosphate buffer, pH 7.2, 2.7% sodium dodecyl sulphate, 1 mM EDTA) at 55 C overnight. Following

hybridization, Northern blots were washed once for 10 minutes in 2.0 X SSPE, 0.1% SDS at room temperature and twice for 10 minutes in 0.15 X SSPE, 0.1% SDS. Autoradiography was carried out from one to three days using Hyperfilm (Amersham) film at -70 C.

Analysis of the levels of RNA levels of HIP1 by Northern blot data revealed that the 10 kilo base HIP1 message is present in all tissue assessed. However, the levels of RNA are not uniform, with brain having highest levels of expression and peripheral tissues having less message. No apparent differences in RNA expression was noted between control samples and HD affected individuals.

EXAMPLE 7

TISSUE LOCALIZATION OF HIP1

Tissue localization of HIP1 was studied using a variety of techniques as described below. Subcellular distribution of HIP-1 protein in adult human and mouse brain Biochemical fractionation studies revealed the HIP1 protein was found to be a membrane-associated protein. No immunoreactivity was seen by Western blotting in cytosolic fractions, using the anti-HIP1-pep1 polyclonal antibody. HIP1 immunoreactivity was observed in all membrane fractions including nuclei (P1), mitochondria and synaptosomes (P2), microsomes and plasma membranes (P3). The P3 fraction contained the most HIP1 compared to other membrane fractions. HIP1 could be removed from membranes by high salt (0.5M NaCl) buffers indicating it is not an integral membrane protein, however, since low salt (0.1- 0.25M NaCl) was only able to partially remove HIP1 from membranes, its membrane association is relatively strong. The extraction of P3 membranes with the non-ionic detergent, Triton X-100 revealed HIP1 to be a Triton X-100 insoluble protein. This characteristic is shared by many cytoskeletal and cytoskeletal-associated membrane proteins including actin, which was used as a control in this study. The biochemical characteristics of HIP1 described were found to be identical in mouse and human brain and was the same for both forms of the protein (both bands of the HIP1 doublet). HIP1 co-localized with huntingtin in the P2 and P3 membrane fractions, including the high-salt membrane extractions, as well as in the Triton X-100 insoluble residue. The subcellular distribution of HIP1 was unaffected by the

expression of polyglutamine-expanded huntingtin in transgenic mice and HD patient brain samples.

The localization of HIP1 protein was further investigated by immunohistochemistry in normal adult mouse brain tissue. Immunoreactivity was seen in a patchy, reticular pattern in the cytoplasm, appeared excluded from the nucleus and stained most intensely in a discontinuous pattern at the membrane. These results are consistent with the association of HIP1 with the cytoskeletal matrix and further indicate an enrichment of HIP1 at plasma membranes. Immunoreactivity occurred in all regions of the brain, including cortex, striatum, cerebellum and brainstem, but appeared most strongly in neurons and especially in cortical neurons. As described previously, huntingtin immunoreactivity was seen exclusively and uniformly in the cytosol.

The in situ hybridization studies showed HIP1 mRNA to be ubiquitously and generally expressed throughout the brain. This data is consistent with the immunohistochemical results and was identical to the distribution pattern of huntingtin mRNA in transgenic mouse brains expressing full-length human huntingtin.

Protein Preparation And Western Blotting For Expression Studies

Frozen human tissues were homogenized using a Polytron in a buffer containing 0.25M sucrose, 20mM Tris-HCl (pH 7.5), 10mM EGTA, 2mM EDTA supplemented with 10ug/ml of leupeptin, soybean trypsin inhibitor and 1mM PMSF, then centrifuged at 4,000rpm for 10' at 4 C to remove cellular debris. 100-150ug/lane of protein was separated on 8% SDS-PAGE mini-gels and then transferred to PVDF membranes. Huntingtin and HIP1 were electroblotted overnight in Towbin's transfer buffer (25 mM Tris-HCl, 0.192M glycine, pH8.3, 10% methanol) at 30V onto PVDF membranes (Immobilon-P, Millipore) as described (Towbin et al, *Proc. Nat'l Acad. Sci.(USA)* 76: 4350-4354 (1979)). Membranes were blocked for 1 hour at room temperature in 5% skim milk/ TBS (10mM Tris-HCl, 0.15M NaCl, pH7.5). Antibodies against huntingtin (pAb BKP1, 1:500), actin (mAb A-4700, Sigma, 1:500) or HIP1 (pAb HIP-pep1, 1:200) were added to blocking solution for 1 hour at room temperature. After 3 x 10 minutes washes in TBS-T (0.05% Tween-20/TBS), secondary Ab (horseradish peroxidase conjugated IgG, Biorad) was applied in blocking solution for 1 hour

at room temperature. Membranes were washed and then incubated in chemiluminescent ECL solution and visualized using Hyperfilm-ECL film (Amersham).

Generation of Antibodies

5 The generation of huntingtin specific antibodies GHM1 and BKP1 is described elsewhere (Kalchman, et al., *J. Biol. Chem.* 271: 19385-19394 (1996)). The HIP1 peptide (VLEKDDLMDMDASQQN, a.a. 76-91 of Seq. ID No. 2) was synthesized with Cys on the N-terminus for the coupling, and coupled to Keyhole limpet hemocyanin (KLH) (Pierce) with succinimidyl 4-(N-maleimidomethyl) cyclohexane-1-carboxylate (Pierce). Female
10 New Zealand White rabbits were injected with HIP1 peptide-KLH and Freund's adjuvant. Antibodies against the HIP1 peptide were purified from rabbit sera using affinity column with low pH elution. Affinity column was made by incubation of HIP1 peptide with activated thio-Sepharose (Pharmacia).

Western blotting of various peripheral and brain tissues were consistent with the RNA
15 data. The HIP1 protein levels observed was not equivalent in all tissues. The protein expression is predominant in brain tissue, with highest amounts seen in the cortex and lower levels seen in the cerebellum and caudate and putamen.

More regio-specific analysis of HIP1 expression in the brain revealed no differential expression pattern in affected individuals when compared to normal controls, with highest
20 levels of expression seen in both controls and HD patients in the cortical regions.

EXAMPLE 8

CO-IMMUNOPRECIPITATION OF HIP1 WITH HUNTINGTIN

Confirmation of the HD-HIP1 interaction was performed using coimmunoprecipitation as follows. Control human brain (frontal cortex) lysate was prepared in the same manner as
25 for subcellular localization study. Prior to immunoprecipitation, tissue lysate was centrifuged at 5000 rpm for 2 minutes at 4 C, then the supernatant was pre-cleared by the incubated with excess amount of Protein A-Sepharose for 30 minutes at 4°C, and centrifuged at the same condition. Fifty microlitres of supernatant (500 mg protein) was incubated with or without antibodies (10 ug of anti-huntingtin GHM1 (Kalchman, et al. 1996)
30 or anti-synaptobrevin antibody) in the total 500 ul of incubation buffer (20mM Tris-Cl

(pH7.5), 40mM NaCl, 1mM MgCl₂) for 1 hour at 4°C. Twenty microlitres of Protein A-Sepharose (1:1 suspension, for GHM1 and no antibody control) or Protein G-Sepharose (for anti-synaptobrevin antibody; Pharmacia) was added and incubated for 1 hour at 4°C. The beads were washed with washing buffer (incubation buffer containing 0.5 % Triton X-100) three times. The samples on the beads were separated using SDS-PAGE (7.5% acrylamide) and transferred to PVDF membrane (Immobilon-P, Millipore). The membrane was cut at about 150 kDa after transfer for Western blotting (as described above). The upper piece was probed with anti-huntingtin BKP1 (1/1000) and lower piece with anti-HIP1 antibody (1/300).

The results showed that when an anti-HIP1 polyclonal antibody was immunoreacted against a blot containing the GHM1 immunoprecipitates from the brain lysate a doublet was observed at approximately 100 kDa. When GHM1 was immunoreacted against the same immunoprecipitate the 350 kDa HD protein was also seen. The specificity of the HD-HIP1 interaction is seen as no immunoreactive bands seen are as a result of the proteins adsorbing to the Protein-A-Sepharose (Lysate + No Antibody) or when a random, non related antibody (Lysate + anti-Synaptobrevin) is used as the immunoprecipitating antibody.

EXAMPLE 9

Subcellular fractionation of brain tissue

Cortical tissue (20-100 mg/ml) was homogenized, on ice, in a 2 ml pyrex-teflon IKA-RW15 homogenizer (Tekmar Company) in a buffer containing 0.303M sucrose, 20mM Tris-HCl pH 6.9, 1mM MgCl₂, 0.5mM EDTA, 1mM PMSF, 1mM leupeptin, soybean trypsin inhibitor and 1mM benzamidine (Wood et al., *Human Molec. Genet.* 5: 481-487 (1996)).

Crude membrane vesicles were isolated by two cycles of a three-step differential centrifugation protocol in a Beckman TLA 120.2 rotor at 4°C based on the methods of Wood et al (1996). The first step precipitated cellular debris and nuclei from tissue homogenates for 5 minutes at 1300 x g (P1). The 1300 x g supernatant was subsequently centrifuged for 20 minutes at 14 000 x g to isolate synaptosomes and mitochondria (P2). Finally, microsomal

and plasma membrane vesicles were collected by a 35 minute centrifugation at 142 000 x g (P3). The remaining supernatant was defined as the cytosolic fraction.

High salt extraction of membranes

5 Aliquots of P3 membranes were twice suspended at 2mg/ ml in 0.5M NaCl, 10mM Tris-HCl, 2mM MgCl₂, pH7.2, containing protease inhibitors (see above). The same buffer without NaCl was used as a control. The membrane suspensions were incubated on ice for 30 minutes and then centrifuged at 142 000 x g for 30 minutes.

10 Extraction of cytoskeletal and cytoskeletal-associated proteins.

To extract cytoskeletal proteins, crude membrane vesicles from the P3 fraction membrane were suspended in a volume of Triton X-100 extraction buffer to give a protein: detergent ratio of 5:1. The composition of the Triton X-100 extraction buffer was based on the methods of Arai et al., *J. Neuroscience* 38: 348-357 (1994) and contained 2% Triton
15 X-100, 10mM Tris-HCl, 2mM MgCl₂, 1mM leupeptin, soybean trypsin inhibitor, PMSF and benzamidine. Membrane pellets were suspended by hand with a round-bottom teflon pestle, and placed on ice for 40 minutes. Insoluble cytoskeletal matrices were precipitated for 35 minutes at 142 000 x g in a Beckman TLA 120.2 rotor. The supernatant was defined as non-cytoskeletal-associated membrane or membrane--associated protein and was removed.
20 The remaining pellet was extracted with Triton X-100 a second time using the same conditions. We defined the final pellet as cytoskeletal and cytoskeletal-associated protein.

Solubilization of protein and analysis by SDS-PAGE and Western Blotting

Membrane and cytoskeletal protein was solubilized in a minimum volume of 1%
25 SDS, 3M urea, 0.1mM dithiothreitol in TBS buffer and sonicated. Protein concentration was determined using the BioRad DC Protein assay and samples were diluted at least 1 X with 5 X sample buffer (250mM Tris-HCl pH 6.8, 10% SDS, 25% glycerol, 0.02% bromophenol blue and 7% 2-mercaptoethanol) and were loaded on 7.5% SDS-PAGE gels (Bio-Rad Mini-PROTEIN II Cell system) without boiling. Western blotting was performed as
30 described above.

Immunohistochemistry

Brain tissue was obtained from a normal C57BL/6 adult (6 months old) male mouse sacrificed with chloroform then perfusion-fixed with 4% v/v paraformaldehyde/0.01 M phosphate buffer (4% PFA). The brain tissues were removed, immersion fixed in 4% PFA for 1 day, washed in 0.01M phosphate buffered saline, pH 7.2 (PBS) for 2 days, and then equilibrated in 25% w/v sucrose PBS for 1 week. The samples were then snap-frozen in Tissue Tek molds by isopentane cooled in liquid nitrogen. After warming to -20 C, frozen blocks derived from frontal cortex, caudate/putamen, cerebellum and brainstem were cut into 14 mm sections for immunohistochemistry. Following washing in PBS, the tissue sections were blocked using 2.5% v/v normal goat serum for 1 hour at room temperature. Primary antibodies diluted with PBS were applied to sections overnight at 4 C. Optimal dilutions for the polyclonal antibodies BKP1 and HIP1 were 1:50. Using washes of 3 x 5 minutes in PBS at room temperature, sections were sequentially incubated with biotinylated secondary antibody and then an avidin-biotin complex reagent (Vecta Stain ABC Kit, Vector) for 60 minutes each at room temperature. Color was developed using 3-3'-diaminobenzidine tetrahydrochloride and ammonium nickel sulfate.

For controls, sections were treated as described above except that HIP1 antibody aliquots were preabsorbed with an excess of HIP1 peptide as well as a peptide unrelated to HIP1 prior to incubation with the tissue sections.

In situ hybridization

In situ hybridization was performed as previously described with some modification (Suzuki et al, *BBRC* 219: 708-713 (1996)). The RNA probes were prepared using the plasmid gt149 (Lin, B., et al., *Human Molec. Genet.* 2: 1541-1545 (1994)) or a 558 subclone of HIP1. The anti-sense and sense single-stranded RNA probes were synthesized using T3 and T7 RNA polymerases and the In Vitro Transcription Kit (Clontech) with the addition of [α^{35} S]-CTP (Amersham) to the reaction mixture. Sense RNA probes were used as negative controls. For HIP1 studies normal C57BL/6 mice were used. Huntingtin probes were tested on two different transgenic mouse strains expressing full-length huntingtin, cDNA HD10366 (44CAG) C57BL/6 mice and YAC HD10366(18CAG) FVB/N mice. Frozen brain sections

(10um thick) were placed onto silane-coated slides under RNase-free conditions. The hybridization solution contained 40% w/v formamide, 0.02M Tris-HCl (pH 8.0), 0.005M EDTA, 0.3 M NaCl, 0.01M sodium phosphate (pH 7.0), 1x Denhardt's solution, 10% w/v dextran sulfate (pH 7.0), 0.2% w/v sarcosyl, yeast tRNA (500mg/ml) and salmon sperm DNA (200mg/ml). The radiolabelled RNA probe was added to the hybridization solution to give 1 x 10⁶ cpm/200 ul/ section. Sections were covered with hybridization solution and incubated on formamide paper at 65 C for 18 hours. After hybridization, the slides were washed for 30 minutes sequentially with 2x SSC, 1x SSC and high stringency wash solution (50% formamide, 2x SSC and 0.1M dithiothreitol) at 65 C, followed by treatment with RNase A (1mg/ml) at 37 C for 30 minutes, then washed again and air-dried. The slides were first exposed on autoradiographic film (b-max, Amersham, UK) for 48 hours and developed for 4 minutes in Kodak D-19 followed by a 5 minute fixation in Fuji-fix. For longer exposures, the slides were dipped in autoradiographic emulsion (50% w/v in distilled water, NR-2, Konica, Japan), air-dried and exposed for 20 days at 4 C then developed as described. Sections were counterstained with methyl green or Giemsa solutions.

EXAMPLE 10

We determined a more precise location of the HIP1 gene on chromosome 7 in the context of a physical and genetic map of chromosome 7, and determined its genomic organization. HIP1 maps by FISH and RH mapping to chromosome band 7q11.23, which contains the chromosomal region commonly deleted in Williams-Beuren syndrome (WS). We used several methods to refine the mapping of HIP1 in this region. PCR screening of a chromosome 7-YAC-library (Scherer et al., *mammalian Genome* 3: 179-181 (1992)) with primers from the 3' UTR of HIP1 resulted in the identification of only a single positive YAC clone (HSC7E512). This YAC clone had previously been shown to map near the Williams syndrome commonly deleted region (Osborne et al., *Genomics* 45: 402-406 (1997)). The HIP1 cDNA was then used to screen a chromosome 7 specific cosmid library from the Lawrence Livermore National Laboratory (LL07NC01), and the RPCI genomic P1 derived artificial chromosome (PAC) library (Pieter de Jong, Rosswell Park, Buffalo, NY). Several PAC and cosmid clones that were already part of pre-assembled contigs in the Williams

syndrome region at 7q11.23 were identified (Fig 5). Restriction enzyme digestion, blot hybridization experiments and PCR screening confirmed that the clones contained the HIP1 gene.

We determined the exon-intron boundaries and intron sizes of HIP1. Primers were
5 designed based on the sequence of the HIP1 transcript and used to sequence directly from the
cosmid, PAC clone and long PCR products from PAC or genomic DNA. Whenever a PCR
fragment generated was longer than predicted from the cDNA sequence, it was assumed to
contain an intron. The size of the introns was determined by sequencing the intron directly
or by PCR amplification of the introns from both genomic DNA and the cosmid or PAC
10 clone from the region. Three sets of overlapping cosmids and a PAC clone that contain the
entire coding sequence of HIP1 were characterized (Fig 5). Cosmid 181G10 and 250F2
were digested with EcoRI and cloned into the plasmid bluescript. Further sequences were
generated from these plasmid subclones. Intron-exon boundary sequences were then
identified by comparing HIP1 genomic and transcript sequence. The gene is contained within
15 75 kb and comprises 29 exons and 28 introns. The intron-exon boundary sequences are
shown in Table 4, along with the exon and intron sizes. A graphic summary of these data is
also shown in Fig. 5. Exons 1 to 28 contained the coding regions. The last and largest
exon of the HIP1 gene was found to contain approximately 7 kb. Most of the intron-exon
junctions followed the canonical GT-AG rule. An AT was found at the 3' splice site of exon
20 1 and an AC at the 5' splice site of exon 2. Sequence data from all the exon-intron borders
of the coding region and 3'-UTR is set forth in Seq. ID Nos. 16-44. (These sequence have
been deposited with GenBank as Accession Nos. AF052261 to AF052288).

Sequence analysis of previously published 5' untranslated region (GenBank accession
U79734) revealed the possibility that the open reading frame extends upstream of the ATG in
25 the exon 4 to a 5' ATG in exon 1. Although we failed to obtain any additional 5' sequences
despite repeated 5' RACE analyses, an additional ATG, 284 bp upstream of the previously
published exon 1 is in the same reading frame and has the surrounding sequence of
TGCCATGTT which is similar to the AGCCATGGG, the consensus Kozak sequence
(Kozak, M. *Nucl. Acids Res.* 15: 8125-8148 (1987)). If translated from this ATG, the protein
30 would be highly homologous to the N-terminal portion of ZK370.3 and yeast Sla2 protein

(Fig. 6). The translated protein in the region of exons 1 to 3 shows an identity of >40% and similarity of >60% to the N-terminal part of ZK370.3. This suggests that the exons 1 to 3 are probably translated.

In western blot studies, HIP1 is identified as a 120 kd protein (11, 23), while the putative translation of the previously published cDNA gives a protein product of estimated molecular weight of approximately 100 kd. If HIP1 gene were translated from the ATG 284 bp upstream of the exon 1, the expected product would have an estimated molecular weight of 122 kd. RNA PCR studies with primers downstream of this ATG and primers in exon 7 amplify expected products of 576 and 600 bp. Taken together these data support the contention that exon 1 extends further 5' and that HIP1 gene is translated from the ATG in exon 1. Sequence analyses showed no TATA, CAAT box or any GC rich promoter sequence upstream of exon 1 ATG. The promoter prediction programs provided by the server <http://dot.imgen.bcm.tmc.edu:9331/seq.search/gene.search.html> did not predict any promoter upstream of the ATG at position -284, (position 0 corresponds to the first nucleotide of published cDNA, GenBank accession U79734). This suggests that HIP1 may have additional exons.

Finally, we evaluated HIP1 gene as a candidate gene for Huntington disease in families without CAG expansion. In a large study of 1022 patients with a clinical diagnosis of HD, no CAG repeat expansion was found in 12 patients who might represent phenocopies of HD. In at least three families, linkage studies have excluded the HD locus at 4p. Mutation in an interacting protein could result in a similar phenotype as illustrated by the discovery of mutations in dystrophin associated proteins in muscular dystrophies. A mutation in HIP1 may result in altered interaction of huntingtin and HIP1 and lead to cellular toxicity as a result of more HIP1 being free in the cytosol. Thus mutations in huntingtin interacting proteins genes may cause a phenotype suggestive of HD. We studied two of the larger families diagnosed with HD without CAG expansion in HD gene, with the highly informative marker D71816 which maps centromeric and very close to HIP1 gene. The clinical findings in both the families were compatible with a diagnosis of HD, although there were atypical features. In family 1733, HIP1 locus appears to be excluded, as there are two recombinants with the marker. Individuals II-5 and II-7 who do not share the haplotype with

the affected individuals are now 41 and 39 years old and have normal neurological examinations.

In the family 1602, a lod score of 1.92 is obtained with the marker D7S1816 at $\theta_{\max}=0$. Sequencing of all the coding exons did not reveal any mutation in any exon sequence. The promoter sequence has not been examined. Subsequently a whole genome scan revealed a higher lod scores for markers on chromosome 20p.

EXAMPLE 11

A mouse brain lambda ZAPII cDNA library (Stratagene # 93609) was screened with various mouse ESTs which showed homology to the human HIP1 cDNA sequence (see Fig. 7). The ESTs were initially isolated from the non-redundant Database of GenBank EST Division by performing a BLASTN using a fragment of the human HIP1 cDNA as the query. We obtained 4 different ESTs which showed homology to HIP1: 1) aa110840 (clone 520282) which is 399bp and shows 58% identity, at the nucleotide level, to position 1880 to 2259 of the HIP1 cDNA. 2) w82687 (clone 404331) which is 420bp and shows 66% identity, at the nucleotide level, to position 2750 to 2915 of the HIP1 cDNA. 3) aa138903 (clone 586510) which is 509bp and shows 88% identity, at the nucleotide level, to position 2763 to 2832 of the HIP1 cDNA. 4) aa388714 (569088) which is 404bp and shows 88% identity, at the nucleotide level, to position 2475 to 2692 of the HIP1 cDNA.

mHIP1:

Fifty nanograms of a 362bp KpnI & PvuII fragment of clone 569088 (containing EST aa388714) was radioactively labeled with [32-P]-dCTP using random-priming. The probe was allowed to hybridize to filters containing $> 2 \times 10^5$ pfu/ml of the mouse brain lambda ZAPII cDNA library (Stratagene # 93609) overnight at 65°C in Church buffer (0.5M sodium phosphate buffer (pH 7.2), 2.7% SDS, 1mM EDTA). The filters were washed at room temperature for 15 minutes with 2XSSPE, 0.1% SDS, then at 65°C for 20 minutes with 1XSSPE, 0.1%SDS and finally twice at 65°C with 0.5 XSSPE, 0.1%SDS. The filters were exposed to X-ray film (Kodak, XAR5) overnight at -70 C. Primary positives were isolated, replated and subsequent secondary positives were hybridized and washed as for the primary

screen. The resulting positive phage was converted into plasmid DNA by conventional methods (Stratagene) and the cDNA termed 4n-n1, was isolated and sequenced 551bp and 541bp from the T7 and T3 end, respectively. 4n-n1 is 2.2kb in length and the T7 end showed 72% identity, at the nucleotide level, to position 1486 to 1715 of the HIP1 cDNA. The 2.2kb insert from 4n-n1 was excised using EcoR1. Fifty nanograms of the 2.2kb insert was used to produced a radioactive probe and used to screen the mouse brain lambda ZAPII cDNA library (Stratagene # 93609) in the same manner as above. The resulting positive phage was converted into plasmid DNA by conventional methods (Stratagene) and the cDNA termed mHIP1a, was isolated and completely sequenced. mHIP1 is 2.3kb in length and showed 85% identity, at the nucleotide level, to position 726 to 3072 of the HIP1 cDNA.

mHIP1a:

Fifty nanograms of a 1.3kb EcoRI & NcoI fragment of clone 404331 (containing EST w82687) was radioactively labeled with [32-P]-dCTP using random--priming. The probe was allowed to hybridize to filters containing $> 2 \times 10^5$ pfu/ml of the mouse brain lambda ZAPII cDNA library (Stratagene # 93609) overnight at 65°C in Church buffer (see above). The filters were washed at room temperature for 15 minutes with 2XSSPE, 0.1% SDS, then at 65°C for 20 minutes with 1XSSPE, 0.1%SDS and finally twice at 65°C with 0.2XSSPE, 0.1%SDS. The filters were exposed to X-ray film (Kodak, XAR5) overnight at -70°C. Primary positives were isolated, replated and subsequent secondary positives were hybridized and washed as for the primary screen. The resulting positive phage was converted into plasmid DNA by conventional methods (Stratagene) and the cDNA termed mHIP1a, was isolated and completely sequenced. mHIP1a is 3.96 kb in length and shows 60% identity, at the nucleotide level, to position 12 to 2703 of the HIP1 cDNA.

EXAMPLE 12

HIP1a:

The entire mHIP1a cDNA sequence was used to screen the non-redundant Database of GenBank EST Division. We identified a human EST, T08283, which showed homology to

mHIP1a. T08383 (clone HIBBB80) is 391bp and shows 87% identity, at the nucleotide level, to position 2904 to 3113 of the mHIP1a cDNA.

5 Fifty nanograms of a 1.6kb HindIII & NotI fragment of clone 404331 (containing EST T08283) was radioactively labeled with [32-P]-dCTP using random-priming. The probe was allowed to hybridize to filters containing > 2x 10⁵ pfu/ml of a human frontal cortex
10 lambda cDNA library overnight at 65 C in Church buffer (see above). The filters were washed at 65 C for 10 minutes with 1XSSPE, 0.1% SDS, and then for 30 minutes and 15 minutes with 0.1XSSPE, 0.1%SDS. The filters were exposed to X-ray film (Kodak, XAR5) overnight at -70 C. Primary positives were isolated, replated and subsequent secondary
15 positives were hybridized and washed as for the primary screen. The resulting positive phage was converted into plasmid DNA by conventional methods (Stratagene) and the cDNA termed HIP1a, was isolated and completely sequenced. HIP1a is 3.2 kb in length and shows 53% identity, at the nucleotide level, to position 876 to 3058 of the HIP1 cDNA.

15

EXAMPLE 13

Following the identification of a 1.2 kb partial human HIP-1 cDNA by yeast two-hybrid interaction studies, a 3.9 kb HIP-1 fragment was isolated from a cDNA library, ligated to a 5' RACE product then subcloned into the mammalian expression vector pCI-neo (Promega). This construct, CMV-HIP-1, expresses HIP-1 from the CMV promoter and was
20 used in the cell expression studies described below. Mouse HIP-1a (mHIP-1a) was also subcloned into a CMV driven expression vector for cell culture expression studies.

EXAMPLE 14

25 Huntingtin proteins with expanded polyglutamine tracts can aggregate into large, irregularly shaped deposits in HD brains, transgenic mice and in vitro cell culture. We have shown that in HEK (human embryonic kidney) 293T cells the aggregation of full-length and larger huntingtin fragments occurs after the cells have been exposed to a period of apoptotic stress. In order to assess the consequence of HIP-1 expression in cultured cells, we used huntingtin aggregation as one marker of viability.

Human embryonic kidney cells (HEK 293T) were grown on glass coverslips in Dulbecco's modified Eagle medium (DMEM, Gibco, NY) with 10% fetal bovine serum and antibiotics, in 5% CO₂ at 37°C. The cells were transfected at 30% confluency with the calcium phosphate protocol by mixing Qiagen-prepared DNA (Qiagen, CA) with 2.5 M CaCl₂, then incubating at room temperature for 10 min. 2X HEPES buffer (240 mM NaCl, 3.0 mM Na₂HPO₄, 100 mM HEPES, pH 7.05) was added to the DNA/calcium mixture, incubated at 37°C for 60 sec, then added to the cells. After 12-18 h, the media was removed, the cells were washed and fresh media was added. At 36 h post-transfection, the cells were exposed to an apoptotic stress by treatment with 35 uM tamoxifen (Sigma) for 1 hour, or left untreated, then processed for immunofluorescence. The cells were washed with PBS, fixed in 4% paraformaldehyde/PBS solution for 20 minutes at room temperature then permeabilized in 0.5% Triton X-100/PBS for 5 min. Following three PBS washes, the cells were incubated with anti-huntingtin antibody MAB2166 (Chemicon) (1:2500 dilution) and anti-HIP-1 antibody HIP-1fp (1:100 dilution) in 0.4% BSA/PBS for 1 h at room temperature in a humidified container. The primary antibody was removed, the cells were washed and secondary antibodies conjugated to Texas red or FITC were added at a 1:600-1:800 dilution for 30 min at room temperature. The cells were then washed again, and the coverslips were mounted onto slides with DAPI (4',6'-diamindino-2 phenylindole, Sigma) as a nuclear counter-stain. Immunofluorescence was viewed using a Zeiss (Axioscope) microscope, digitally captured with a CCD camera (Princeton Instrument Inc.) and the images were colourized and overlapped using the Eclipse (Empix Imaging Inc.) software program. Appropriate control experiments were performed to determine the specificity of the antibodies, including secondary antibody only and mock transfected cells.

The huntingtin fragment HD1955 was used in the aggregation studies. This fragment represents the N-terminal 548 amino acids of huntingtin, and corresponds approximately to the polyglutamine-containing fragment produced by caspase 3 cleavage of huntingtin. Transfection of HD1955 with 15 polyglutamines (HD1955-15) results in a diffuse cytoplasmic distribution of the expressed protein. Transfection of HD1955 with 128 polyglutamines (HD1955-128) also results in diffuse cytoplasmic expression. However, exposure of cells transfected with HD1955-128 to tamoxifen results in a marked

redistribution of huntingtin. In 29% of cells expressing HD1955-128, the huntingtin protein appears as dense aggregates that are localized in the perinuclear area of the cell. In contrast, less than 1% of HD1955-128 expressing cells contain aggregates in the absence of tamoxifen, and 0% of HD1955-15 cells form aggregates in the presence or absence of tamoxifen treatment.

Co-transfection of HIP-1 and HD1955 was used to test the influence of HIP-1 on huntingtin aggregation. As a control, b-galactosidase was co-transfected with HD1955. In the control transfections, 1-2% of cells expressing HD1955-128 formed aggregates in the absence of tamoxifen, similar to HD1955-128 expressed alone. However, when HD1955-128 was co-expressed with HIP-1, an average of 14% of huntingtin-expressing cells contained aggregates with no tamoxifen treatment. Double-labeling demonstrated that the majority of the cells containing aggregates also expressed HIP-1, directly implicating HIP-1 in the increase in aggregation. Therefore, these results indicate that HIP-1 provides sufficient stress on the huntingtin-expressing cells to form aggregates, to the extent that tamoxifen is no longer necessary.

EXAMPLE 15

We next designed a series of experiments to identify a region of HIP-1 sufficient for inducing aggregate formation of HD1955-128. As described above, HIP-1 contains a domain with high homology to the death effector domains (DED) present in many apoptosis-related proteins. The DED domain of HIP-1 was ligated in-frame to HD1955-128, 3' from the caspase-3 cleavage site. Transfection of the resulting fusion protein with the DED ligated in the sense orientation (HD1955-128-DEDsense) resulted in a large number (30-50%) of cells containing aggregates, without tamoxifen incubation. In contrast, expression of a huntingtin-DED fusion protein with DED in the antisense orientation (HD1955-128-DEDantisense) did not have more aggregates than the HD1955-128 no tamoxifen control. Therefore, the DED domain of HIP-1 is sufficient to stress the cells, causing aggregate formation.

EXAMPLE 16

To directly assess the effect of HIP-1 expression on cell viability, mitochondrial function tests were performed on 293T cells transfected with HIP-1. The assessment of mitochondrial function, using the MTT assay (Carmichael et al., *Cancer Res.* 47: 936-942 (1987); Vistica et al., *Cancer Res.* 51: 2515-2520 (1991)), is a standard method to measure cell viability. The MTT assay quantitates the formation of a coloured substrate (formazan), with the mitochondria of viable cells forming more substrate than non-viable cells. Since decreased mitochondrial activity is an early consequence of many cellular toxins, the MTT assay provides an early indicator of cell damage.

For cell viability assays, HEK 293T cells were seeded at a density of 5×10^4 cells into 96-well plates and transfected with 0.1 ug or 0.08 ug HIP-1 or 0.1 ug of the control construct lacZ using the calcium phosphate method described above. At 24-36 hours post-transfection tamoxifen-treated cells were incubated for 2 hours in a 1:10 dilution of WST-1 reagent (Boehringer Mannheim) and release of formazan from mitochondria was quantified at 450 nm using an ELISA plate reader (Dynatech Laboratories) (Carmichael et al., 1987; Mosmann, *J. Immunol. Meth* 65: 55-63 (1983)). One way ANOVA and Newman-Keuls test were used for statistical analysis. The transfection efficiency, measured by β -galactosidase staining and immunofluorescence, was approximately 50%.

We have previously demonstrated that expression of mutant huntingtin results in increased susceptibility to an apoptotic stress induced by sub-lethal doses of tamoxifen in transfected 293T cells (Martindale et al., 1998). A similar assay was used to test the consequence of HIP-1 expression. With 0.1 ug transfected HIP-1 DNA, after 24 hr expression, HIP-1 resulted in increased cell death in response to tamoxifen, compared with the tamoxifen-treated β -galactosidase control ($p < 0.01$, $n=4$). Reducing the amount of transfected HIP-1 DNA to 0.08 ug also resulted in increased cell death compared with control ($p < 0.01$, $n=4$), indicating the high potency of HIP-1 (Fig. 8). Furthermore, increased cell death in cells transfected with HIP-1 was observed in the absence of apoptotic stress at 48 hrs post-transfection, but was so severe that it could not be accurately quantitated. Thus, an earlier time point (24 hr) had to be used for better reproducibility, using an apoptotic stress to unmask the phenotype.

In order to model a pathogenic interaction of HIP-1 and huntingtin in the HEK 293 mammalian cell system, HIP-1 was transfected into cell lines stably expressing huntingtin. Two cell lines were chosen for the initial studies, one line expressed the truncated HD1955 construct with 15 glutamines, and the second expressed the HD1955 with 128 repeats. Western blotting indicated that the cell lines expressed huntingtin at similar levels. To assess whether HIP-1 is toxic in the presence of mutant huntingtin, 0.1 ug HIP-1 DNA was transfected into the HD1955-128 cell line. Transfection of HIP-1 into the HD1955-15 cell line was used as the wild-type huntingtin control, and transfection of LacZ into both cell lines was the control for transfection-related toxicity (Figs 9A and 9B). MTT toxicity assays showed that HIP-1 in the presence of mutant huntingtin (HD1955-128) was significantly more toxic than HIP-1 with wild-type huntingtin (HD1955-15), $p < 0.001$, $n = 4$ (Fig. 9C). This toxicity was observed at 24 hr and 36 hr post-transfection. No tamoxifen was needed to unmask the phenotype, suggesting that the combined cell stress of HIP-1 with truncated huntingtin was sufficient to reduce cell viability over control.

CLAIMS

- 1 1. A polypeptide comprising the sequence given by Seq. ID. No. 5.
- 1 2. A cDNA molecule comprising the sequence given by Seq. ID No. 6.
- 1 3. A polypeptide comprising the sequence given by Seq. ID No. 7.
- 1 4. A method for ameliorating the effects of Huntington's disease in a
2 patient expressing a HIP-apoptosis modulating protein, comprising the step of administering
3 the patient a therapeutic composition which reduces the activity of the HIP-apoptosis
4 modulating protein.
- 1 5. A method according to claim 4, wherein the composition comprises a
2 material which binds to HIP-apoptosis modulating protein.
- 1 6. The method according to claim 4, wherein the composition comprises
2 an expression vector encoding huntingtin having a normal number of repeats.
- 1 7. An expression vector for expression of a gene in a mammalian host
2 comprising a region encoding an HD-interacting polypeptide.
- 1 8. The expression vector according to claim 7, wherein the HD-
2 interacting polypeptide is an HIP-apoptosis modulating protein.
- 1 9. The expression vector according to claim 8, wherein the HIP-apoptosis
2 modulating protein has a sequence which includes the amino acid sequences given by SEQ
3 ID Nos. 2, 4, 5 or 7.

1 10. The expression vector of claim 7, wherein the HD-interacting
2 polypeptide interacts differently with expanded Huntingtin than with Huntingtin having a
3 CAG repeat region containing 15 to 35 repeats.

1 11. The expression vector according to claims of claims 7-10, further
2 comprising a region encoding Huntingtin having a polyglutamine tract of 35 or fewer.

1 12. A method for inducing apoptotic death in cells, comprising the step of
2 introducing into the cells an expression vector encoding at least the death effector domain of
3 a HIP-apoptosis modulating protein whereby the death effector domain is expressed by the
4 cells.

1 13. The method of claim 12, wherein the expression vector encodes the
2 amino acid sequence given by Seq. ID. No. 2.

1 14. The method of claim 12, wherein the expression vector encodes the
2 amino acid sequence given by Seq. ID. No. 4.

1 15. A method for screening a composition for the ability to inhibit
2 apoptosis induced by an HIP-apoptosis modulating protein, comprising simultaneously
3 exposing a population of cells to the composition and an HIP-apoptosis modulating protein
4 and measuring the extent of cell death.

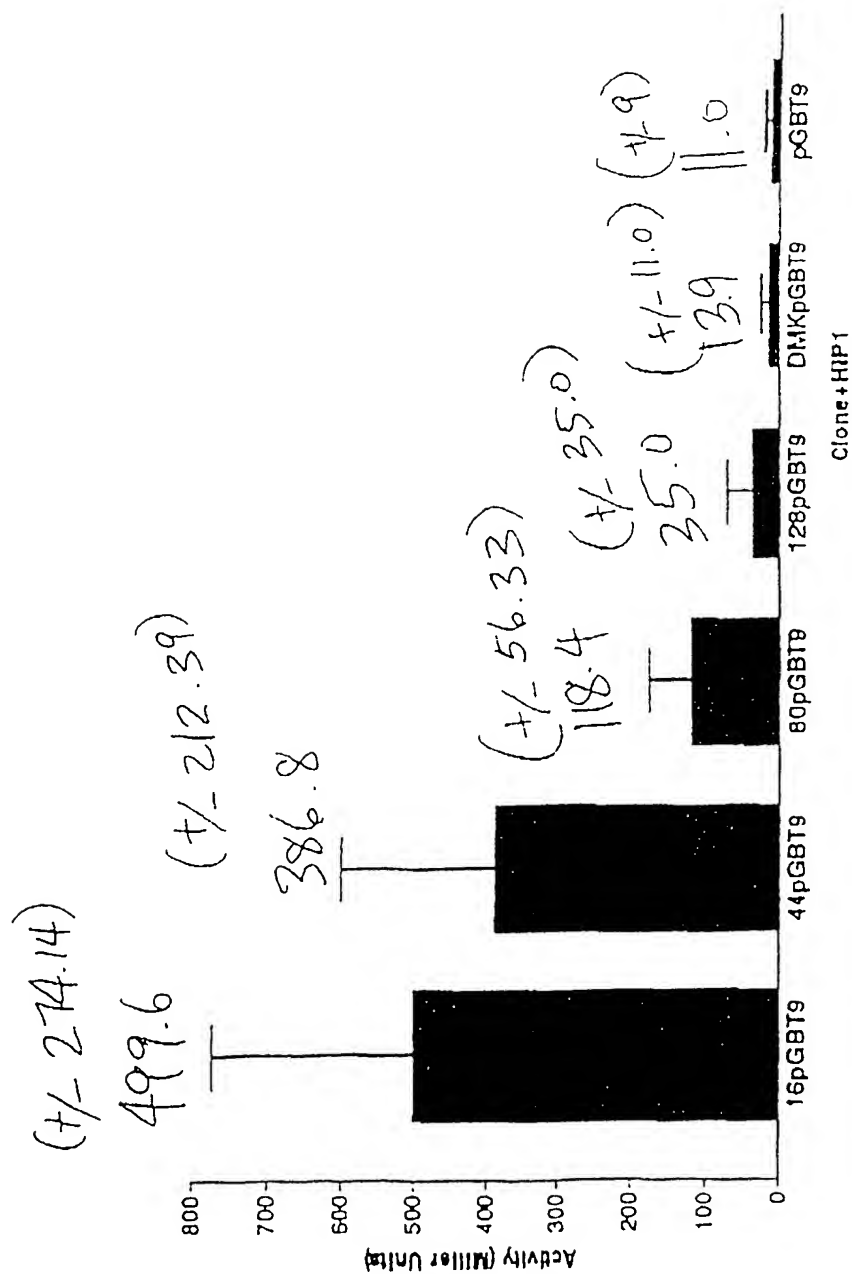


Fig. 1

Fig 2

HIP1 Clones: Nucleotide Alignment

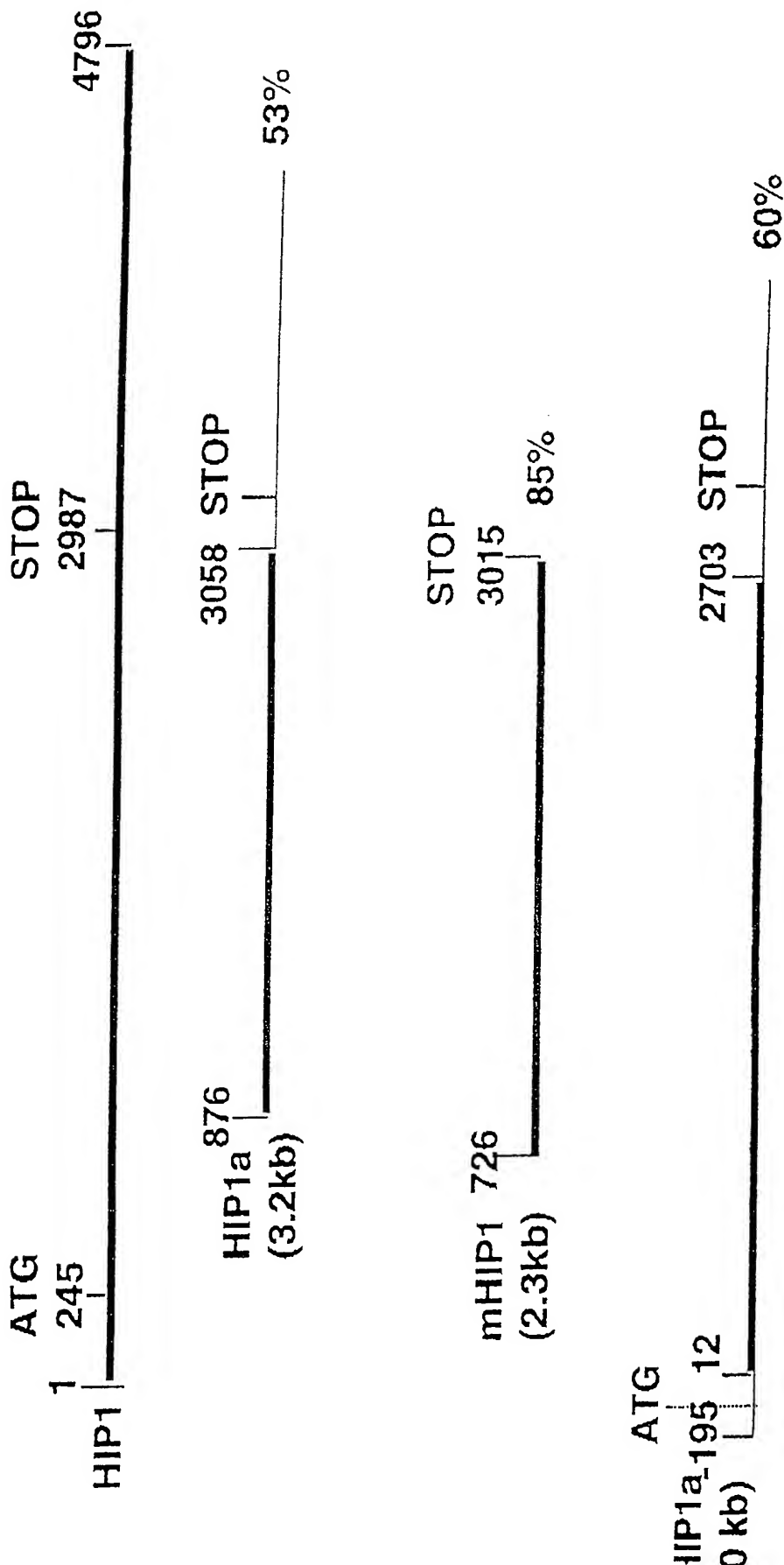


Fig 3 HIP1 Clones: Protein Alignment

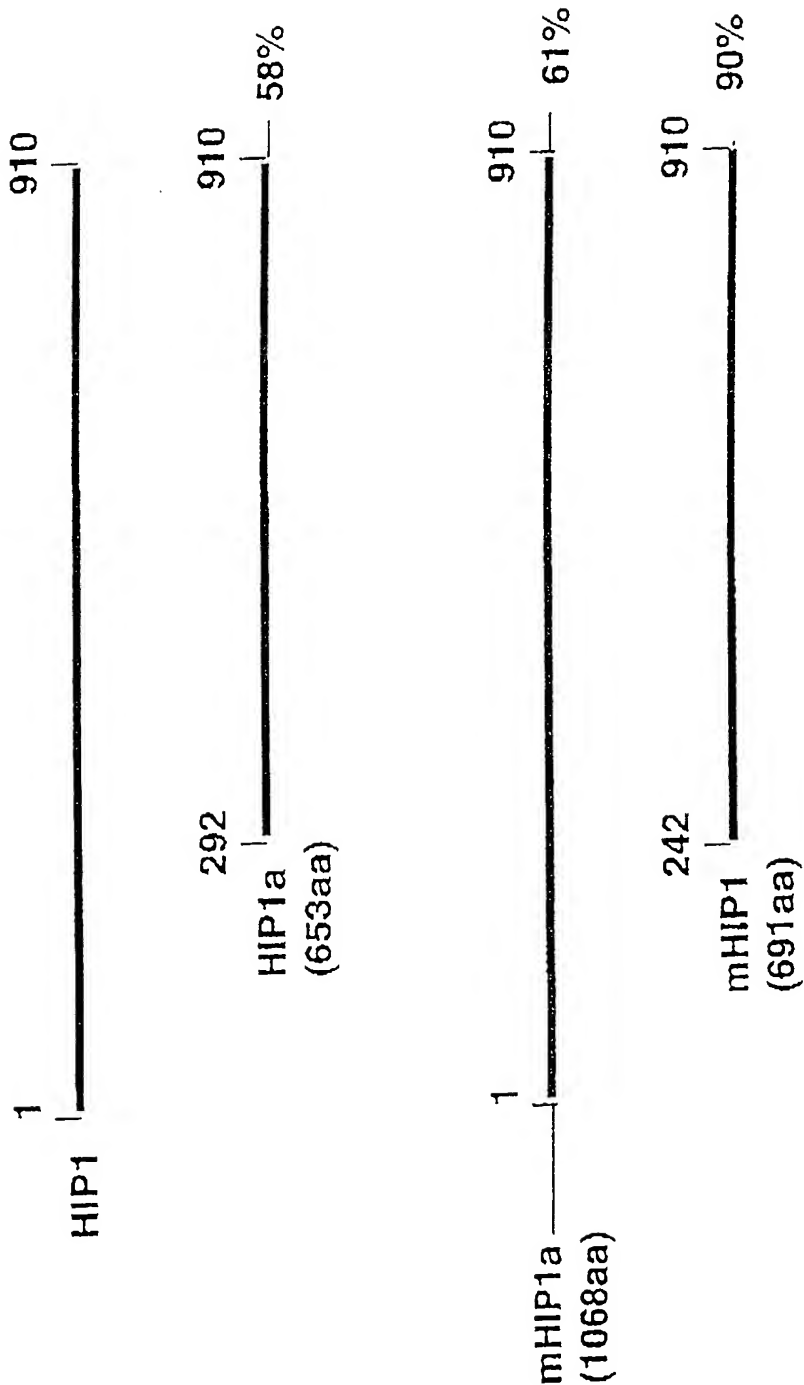


Fig 4

>Usurpin A
SAEVIHQVEEALDTDEKIMLLFLCRDVAIDVVPNVRLLDILRERGKLSVCDLAELLYRVHRFDLLKRILK

>Usurpin B
YRVLMAHIGEDLDKSDVSSLIFLMKDYMGGRGKISKHKSFDDLVLVHKLNLVAPDQLDLEKCLKNIHRIDLKTKIQK

>Casp-8 A
FSRNLVDIGELQDSEDLASLKELSLDYIPQRKOEPIKDALMIFQRLQEKRMLEESNLSFLKELLFRINRLDLLITYLN

>Casp-8 B
YRVMLYQISEEVSREELRSFKFLLQHEISKCKLDDDMNLLDIFIEMEKRVILGEGKLDILKRVCAQINKSLLKIND

>Casp-10 A
FRHKLLTIDSNLGVQDVENLKFCLIGLVPNKKLEKSSGASDVFEHLLANDLLSEEDPFFLAELLYIIRQKKLLQHLNC

>Casp-10 B
FRNLLYELSEGIDSENKDMIFLLKDSLPKTEMTSLSFALFLEKQGGKIDEDNLTCLDLCKTVVPKLLRNIEK

>FADD
FLVLHSHVSSSLSSSELTFLKFLCLGRVGRKLERVQSGLDLFSMLLEQNDLEPGHTELLRELLASLRHDLLRRVDD

>MC159 A
SLPFLRHLLLEELDSHEDSLLLFLGHDAAPGCTTVTQALCSLSQQRKLTALALVEMLYVLQMDLLKSRFG

>MC159 B
YHKLMVCVGEELDSSSELRALRLFACNLNPSLSTALSESSRPVELVLALENVGLVSPSSVSVLADMLRTLRLDLCCQQLVE

>E8
FRCLMALVNDFLSDKEVEEHYFLCAPRLESHLEPGSKKSFRLASLLEDLELLGGDKLTLRHLTTIGRADLVKNLQV

>KS orfk13A
TYEVLCEVARKLGTDDREVVLFLNVFLPQPTLAQIIGALRALKKEGRLTFPLLAECLEPRAGRRDLLRDLLH

>KS orfk13B
YQLTVLHVDGELCARDIRSLIFLSKDTIGSRSTPQTFLNHVYCMENLDLLGPTDVDALMSMLRSLSRVDLQRQVQT

>HIP1
SELEADLAEQQHLRQQAADDCEFLRAELDELRRQREDTEKAQRSLSIEIERKAQANEQRYSKLKEKYSSELVQNHADLLRKN
AE

>HIP1a
GELEEQRKQKQKALVDNEQLRHELAQLRAAQLEERSQGLREEAERKASATEARYNKLKEKHSELVHVHAEELLRKNAD

>mHIP1a
NGLEAELEEQRKQKQKALVDNEQLRHELAQLKALQLEGARNQGLREEAERKASATEARYSKLKEKHSELINTHAEELLRKN
AD

>mHIP1
SELEAELEQQHLGRQAMDDCEFLRTELDELKRQREDTEKAQRSLSIEIERKAQANEQRYSKLKEKYSSELVQNHADLLRKN
AE

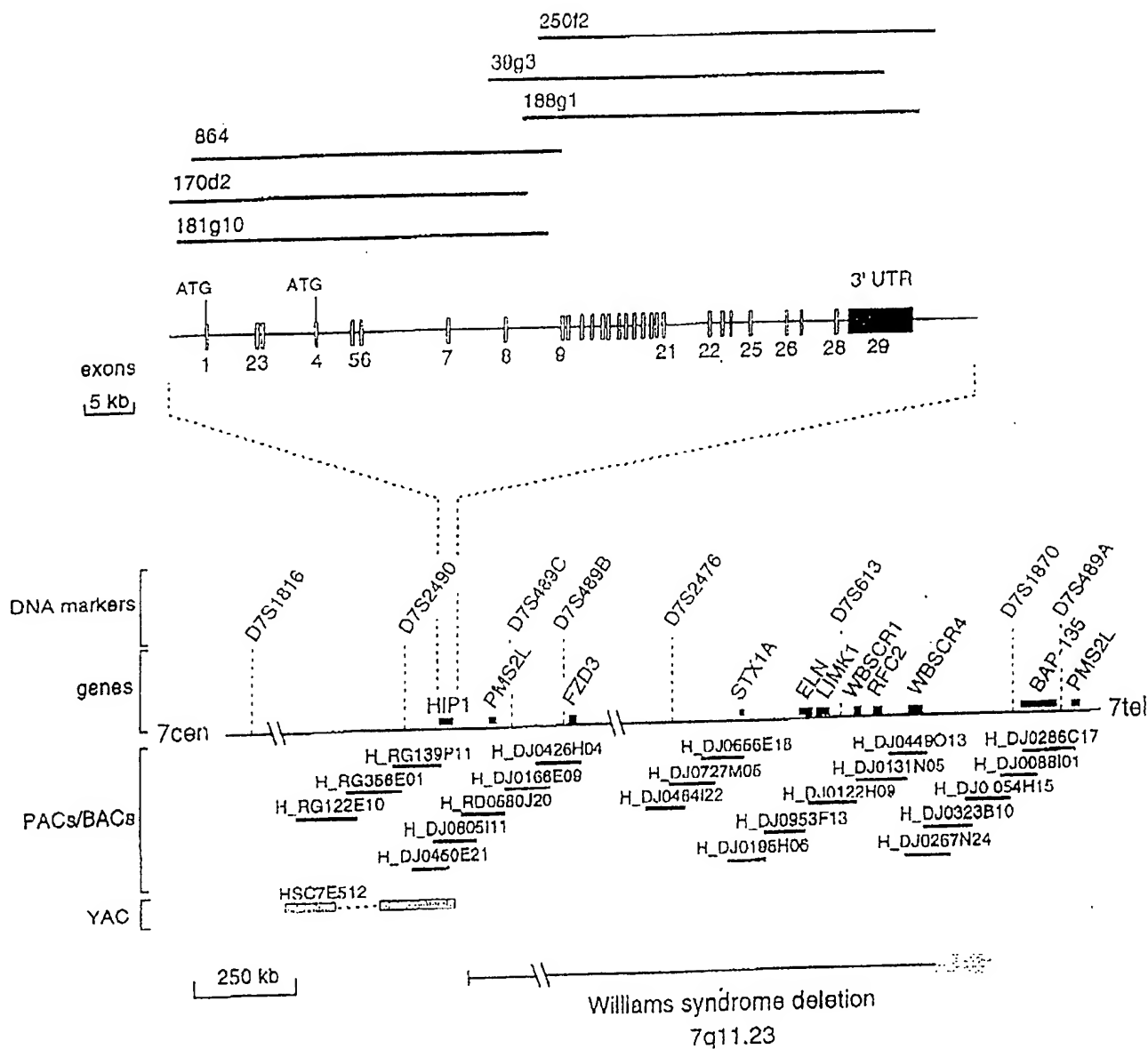


Fig 5

Fig 6

1 hip1 MLCQSEWRDQQL GTANARQWCEPLPQDA QPAGSHERCEPPIPPA GRLOQIDHPGWGRLL AGGGERGLWEGLSH SORLIHLTLLSLPLL 90
2 zk370.3 -----MDHRAQAREVFV 12

1 hip1 Q KAI E . K KHART I.GT E EK FW V L VL KKECH . HCLLDGEP V RY N S 180
2 zk370.3 VQTVSINKAINTQE VAVREKHARTGILGT HHEKGAQTFSVWNR LPLSSNAVLCKEKE VFKLLRDGHPNVLK DSLRYBNELSDMSRM 102
3 BAQLEAVQKAITQNE VPLKPKHARTIIVGT HHEKSSGIFNHTVGR IQLRKHPTLTWKKECH LVHKLLEDGHRKVS ETRYVNRPTQLSQE

1 hip1 W HL GYG Y KLL .H K P PG L D QL E D .H F T M L V. R S. 269
2 zk370.3 WHLTSYGYPGIES YCKLHURVTTERKY PWPQKLDLNDSQL TL-EGDLONFFETI DMLQDALLVLQDR VYENRNSLRNSLIP 191

1 hip1 GQC PLI .ILD S YDY VK .KEL ES . D L G ER RF P K . SSNLQTFK L. IL LP PNEF S 355
2 zk370.3 AGCRLAPLQVILD CSHLYDYTVKILFKL ESCLP ----ADTLQG ERDRFMSQFTKLDL FYRSSNLQTFKRLIQ IPQLPENPPNPLRAS 277
3 QGQMLSPRLAILED TSKFYDYLVMKIFKL ESQVP ----PDALGG HRSRFTIPTERTKF YEESNLQTFKYLVS IPTFESHAPNLTQOS

1 hip1 L P E S D H D D I L . K E 445
2 zk370.3 ALSEHISPVVPIPAE ASSPDSPEVLEKDDG KMDASQNFEDNKF DDIFGSSPSPDPNF NSQNGWIKDKDHLI ERLYREISGLKRALE 346
3 DLESYRTPHAYLHSE GS-----E--- --DGTSLDGHGEL LNLAAEPQ--ASP SSQ-----PDPEEQI VMLSRVDEDEFAKE

1 hip1 E . Q . L E R T A . E ERKA A E R K X Y 530
2 zk370.3 RMKTS---QRVLQ LKGVSEIEADLAEQ QHLRQQRADDCPELR AEIDELRRQREDTEK AQRSLSEIERKAAQAN EQ-RYSKLEKRYSE- 429
3 RLIOEA---RSRIEQ YENRLLQMGFEFDA KREADENREBAQRK NELALRDASRTQTD AR--VKEAEKATNA EE-RFNKMGVTEK.

1 hip1 H L KQ. D L QR S 617
2 zk370.3 ---LVQNEADLERN ARVTQVSHARQAV DLREKKLESDSLER ISDQGRKTOEQLV LBSLQBLATSOREL QVLQSLTSFQSEA 492
3 ---FASBEVLALTKL GDIQKLEASEISKF DRDEZ---ITALNR KVEEAQR-----BAGRATKA EGDAGNVDEMRTOJV

1 hip1 E EL D H E Q AK . E Q A P 702
2 zk370.3 HNAEPALEKERS LVSGAHRRELSAL RKELODTQLKLASTE ESMCQAKDQRHLL VGRKRAEQVIQDAL NQLEPPLIS----- 569
3 KADIEVELEKRTID- -----HURESHAN- -QLVQSSAREBTKIR LAELEVAES-GVGI TOMFDKCEDALQNT SITYP-----

1 hip1 HL I E L A EL S A A CK A 792
2 zk370.3 CAGSACHULSTVTSI SSCIEQLEKSMQYL ACPEDISGLHSITL LAHLTSDAINAGATT CLRAPPADSLTBA CKQYGRETLAYLASL 641
3 -----HLAQSAMU LVNLSNR-LDEPL ATKONY-----F AGHLLSITLSAASA AYTASISSEYGVNDQ CEKV---LAAAKVAF

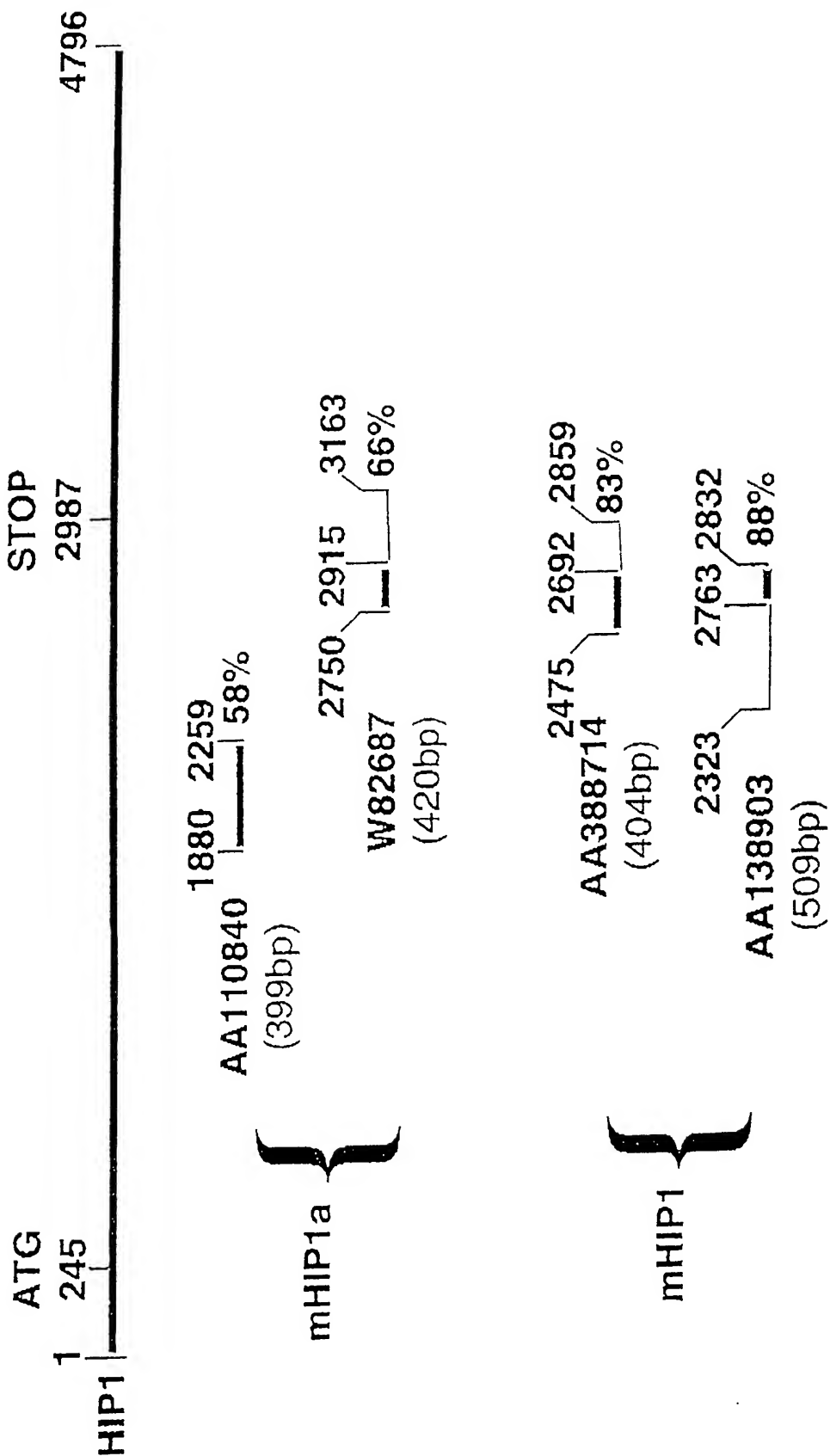
1 hip1 L D LP DI . EM AI A IE RA G LEVNE IL 882
2 zk370.3 BEEGSLENDASTANR NCLSKIKAIGBELLP RGLDIKQELGDLVD KEMAPTSAAITATA RIBEMLSKSRAGDTG VLEWNERILGCCTS 731
3 SDESALSRADKHELL RDIQTLSLSHISLP LQTDIDKDVWNELE QERRRMDAIRFAVQ BIEAIQRRARSSDQ IRLVNESILANCA

Fig 6 (cont.)

1	1	1	L..AS	L	Q	ZIV	SP	KEV	N	MTGLIS	A	KAVG	A	V	V	AD	VV	GRGK	F	K	L	V	BIAASTA	958
2	2	2	370.3	LMQAIQVLI'VASKDL	QREIVESGRGTASPK	EFVAKNSHTBGLIS	ASKAVGHGATVHTDA	ADLVYQGRGK	----	F	KELAVCSHEIAASTA	917												
1	1	1	1	L	A	VNQ	TA	VVA	G	.	D?S	L	K	EX	SQV	L	ELE	L	ER	KL	1056			
2	2	2	370.3	QLVAAASKVKADEDSR	NLAQLQOASRGVHQA	TAGVVASTISGKSQI	BETDM--	MDPSSHTL	TQIKRQEDSQVRVL	ELEHVLQZKQKLE	505													
1	1	1	1	L	A	VNQ	TA	VVA	G	.	D?S	L	K	EX	SQV	L	ELE	L	ER	KL	1056			
2	2	2	370.3	QLEVSSPVRADYDSS	KIDALSVAUKAVHQN	TACVVAVXNGZTTL	NDEGS--	LDPSYLSL	HAAKKEEESQVYOML	BLEQSLNQERAKLAA	505													
1	1	1	1	L	A	VNQ	TA	VVA	G	.	D?S	L	K	EX	SQV	L	ELE	L	ER	KL	1056			
2	2	2	370.3	LRKBY	A	.	LRKBYELAGVABGW	EEGTASPPPTLOEVV	TEKE	1090														
1	1	1	1	L	A	VNQ	TA	VVA	G	.	D?S	L	K	EX	SQV	L	ELE	L	ER	KL	1056			
2	2	2	370.3	LRKBYELAGVABGW	EEGTASPPPTLOEVV	TEKE	1090																	
1	1	1	1	L	A	VNQ	TA	VVA	G	.	D?S	L	K	EX	SQV	L	ELE	L	ER	KL	1056			
2	2	2	370.3	LRKBYELAGVABGW	EEGTASPPPTLOEVV	TEKE	1090																	

Fig 7

Mouse ESTs



mthlp1.pzm:Graph-2 - Tue Apr 28 11:30:41 1998

Hip-1 increases the susceptibility to cell stress

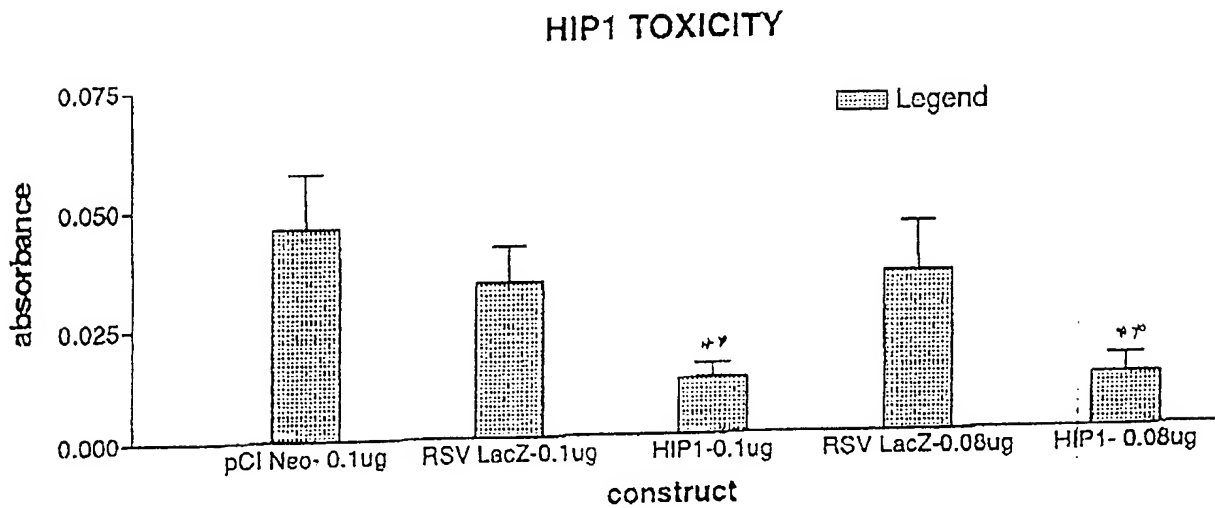


Fig 8

Hip-1 is toxic in the presence of huntingtin

HIP1 transfected into HD1955-15 stable cell line
36 hr post-transfection

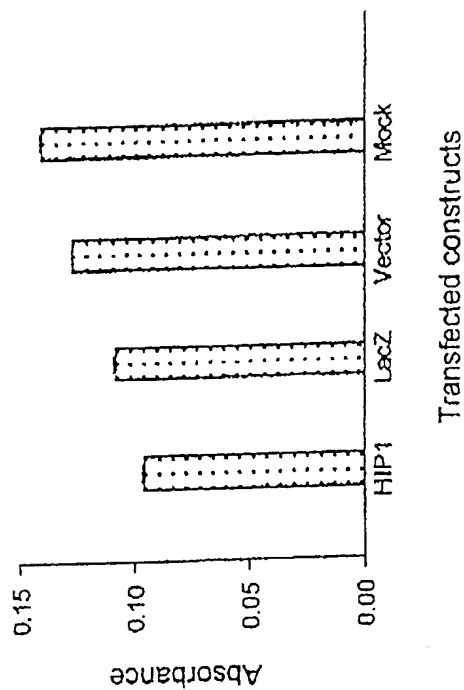


Fig 9A

Hip-1 is toxic in the presence of huntingtin

HIP1 transfected into HD1955-128 stable cell line
36 hr post-transfection

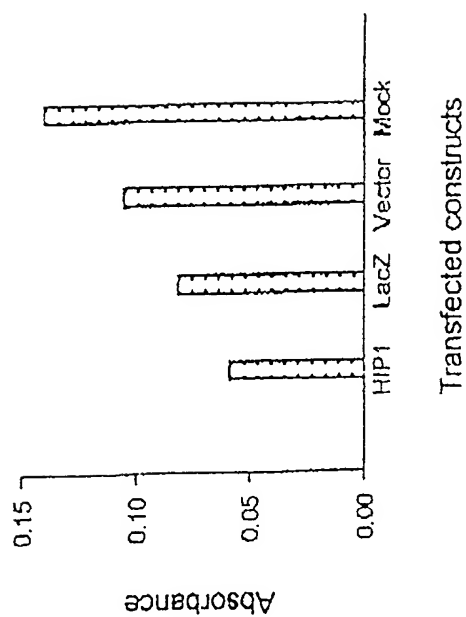
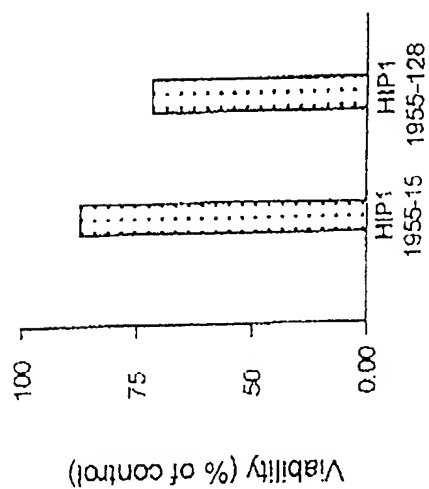


Fig 9B

Polyglutamine-dependence of HIP-1 toxicity



Transfected constructs/cell lines

Fig 9c

SEQUENCE LISTING

(1) GENERAL INFORMATION:

(i) APPLICANT: Kalchman, Michael

Hayden, Michael R.

Hackam, Abigail

Chopra, Vikramjit Singh

Nicholson, Donald W.

Vallaincourt, John P.

Rasper, Dita M.

(ii) TITLE OF INVENTION: Apoptosis Modulators That Interact with the
Huntington's Disease Gene

(iii) NUMBER OF SEQUENCES: 44

(iv) CORRESPONDENCE ADDRESS:

(A) ADDRESSEE: Oppedahl & Larson

(B) STREET: PO Box 5270

(C) CITY: Frisco

(D) STATE: CO

(E) COUNTRY: USA

(F) ZIP: 80443-5270

(v) COMPUTER READABLE FORM:

(A) MEDIUM TYPE: Diskette, 3.50 inch, 1.44 Kb storage

(B) COMPUTER: IBM Compatible

(C) OPERATING SYSTEM: MS DOS 5.0

(D) SOFTWARE: WordPerfect

(vi) CURRENT APPLICATION DATA:

(A) APPLICATION NUMBER:

(B) FILING DATE:

(C) CLASSIFICATION:

(viii) ATTORNEY/AGENT INFORMATION:

(A) NAME: Larson, Marina T.

(B) REGISTRATION NUMBER: 32038

(C) REFERENCE/DOCKET NUMBER: UBC.P-013US2

(ix) TELECOMMUNICATION INFORMATION:

(A) TELEPHONE: (970) 668-2050

(B) TELEFAX: (970) 668-2052

(2) INFORMATION FOR SEQ ID NO:1:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1164

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: cDNA for Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

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TTTGGCAGTT CATCCAGCAG TGATCCCTTC AATTTCACAA GTCAAAATGG 350
TGTGAACAAG GATGAGAAGG ACCACTTAAT TGAGCGACTA TACAGAGAGA 400
TCAGTGGATT GAAGGCACAG CTAGAAAACA TGAAGACTGA GAGCCAGCGG 450
GTTGTGCTGC AGCTGAAGGG CCACGTCAGC GAGCTGGAAG CAGATCTGGC 500
CGAGCAGCAG CACCTGCGGC AGCAGGCGGC CGACGACTGT GAATTCCTGC 550
GGGCAGAACT GGACGAGCTC AGGNGGCAGC GGGAGGACAC CGAGAAGGCT 600
CAGCGGAGCC TGTCTGAGAT AGAAAGGAAA GCTCAAGCCA ATGAACAGCG 650
ATATAGCAAG CTAAAGGAGA AGTACAGCGA GCTGGTTCAG AACCACGCTG 700
ACCTGCTGCG GAAGAATGCA GAGGTGACCA AACAGGTGTC CATGGCCAGA 750
CAAGCCCAGG TAGATTTGGA ACGAGAGAAA AAAGAGCTGG AGGATTTCGT 800
GGAGCGCATC AGTGACCAGG GCCAGCGGAA GACTCAAGAA CAGCTGGAAG 850
TTCTAGAGAG CTTGAAGCAG GAACTTGGCA CAAGCCAACG GGAGCTTCAG 900
GTTCTGCAAG GCAGCCTGGA AACTTCTGCC CAGTCAGAAG CAAACTGGGC 950
AGCCGAGTTC GCCGAGCTAG AGAAGGAGCG GGACAGCCTG GTGAGTGGCG 1000
CAGCTCATAG GGAGGAGGAA TTATCTGCTC TTCGGAAAGA ACTGCAGGAC 1050
ACTCAGCTCA AACTGGCCAG CACAGAGGAA TCTATGTGCC AGCTTGCCAA 1100
AGACCAACGA AAAATGCTTC TGGTGGGGTC CAGGAAGGCT GCGGAGCAGG 1150
TGATACAAGA CGCG 1164
```

(2) INFORMATION FOR SEQ ID NO:2:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 386

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

```
Thr Ala Asp Thr Leu Gln Gly His Arg Asp Arg Phe Met Glu Gln
1           5           10           15
```

```
Phe Thr Lys Leu Lys Asp Leu Phe Tyr Arg Ser Ser Asn Leu Gln
20           25           30
```

```
Tyr Phe Lys Arg Val Ile Gln Ile Pro Gln Leu Pro Glu Asn Pro
```

	35		40		45
Pro Asn Phe Leu Arg Ala Ser Ala Leu Ser Glu His Ile Ser Pro	50		55		60
Val Val Val Ile Pro Ala Glu Ala Ser Ser Pro Asp Ser Glu Pro	65		70		75
Val Leu Glu Lys Asp Asp Leu Met Asp Met Asp Ala Ser Gln Gln	80		85		90
Asn Leu Phe Asp Asn Lys Phe Asp Asp Phe Gly Ser Ser Ser Ser	95		100		105
Ser Asp Pro Phe Asn Phe Asn Ser Gln Asn Gly Val Asn Lys Asp	110		115		120
Glu Lys Asp His Leu Ile Glu Arg Leu Tyr Arg Glu Ile Ser Gly	125		130		135
Leu Lys Ala Gln Leu Glu Asn Met Lys Thr Glu Ser Gln Arg Val	140		145		150
Val Leu Gln Leu Lys Gly His Val Ser Glu Leu Glu Ala Asp Leu	155		160		165
Ala Glu Gln Gln His Leu Arg Gln Gln Ala Ala Asp Asp Cys Glu	170		175		180
Phe Leu Arg Ala Glu Leu Asp Glu Leu Arg Gln Arg Glu Asp Thr	185		190		195
Glu Lys Ala Gln Arg Ser Leu Ser Glu Ile Glu Arg Lys Ala Gln	200		205		210
Ala Asn Glu Gln Arg Tyr Ser Lys Leu Lys Glu Lys Tyr Ser Glu	215		220		225
Leu Val Gln Asn His Ala Asp Leu Leu Arg Lys Asn Ala Glu Val	230		235		240
Thr Lys Gln Val Ser Met Ala Arg Gln Ala Gln Val Asp Leu Glu	245		250		255
Arg Glu Lys Lys Glu Leu Glu Asp Ser Leu Glu Arg Ile Ser Asp	260		265		270
Gln Gly Gln Arg Lys Thr Gln Glu Gln Leu Glu Val Leu Glu Ser	275		280		285
Leu Lys Gln Glu Leu Gly Thr Ser Gln Arg Glu Leu Gln Val Leu					

	290		295		300
Gln Gly Ser Leu	Glu Thr Ser Ala Gln	Ser Glu Ala Asn Trp	Ala		
	305		310		315
Ala Glu Phe Ala	Glu Leu Glu Lys Glu	Arg Asp Ser Leu Val	Ser		
	320		325		330
Gly Ala Ala His	Arg Glu Glu Glu Leu	Ser Ala Leu Arg Lys	Glu		
	335		340		345
Leu Gln Asp Thr	Gln Leu Lys Leu Ala	Ser Thr Glu Glu Ser	Met		
	350		355		360
Cys Gln Leu Ala	Lys Asp Gln Arg Lys	Met Leu Leu Val Gly	Ser		
	365		370		375
Arg Lys Ala Ala	Glu Gln Val Ile Gln	Asp Ala			
	380		385 386		

(2) INFORMATION FOR SEQ ID NO:3:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 4796

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: cDNA for Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:

CAGTGTACGG	TTGATCATAT	AACGCCGCGG	GCGGGGATTG	GTTTATATAT	50
CGCAAATTGA	TNTAGGGGGG	GGGGGATGGN	CAGAGATTTC	GCTTCATTAG	100
GCCATTATAA	GCAGGAAGGG	TTTCAAGGAA	AAAAACCCAG	AAAGTGCATA	150
TTGCACCCAC	CATGAGAAAG	GGGCAACAGA	CCTTNTGTTN	TGTTNTCAAC	200
CGCCTGCTTC	TGTTTTAGCA	ACGCAGTGTT	TTGGTGGAAG	TTGTGCCATG	250
TGTTCCACAA	ANTCTTCCGA	GATGGACACC	CGAACGTCCT	GAAGGACTTT	300
GTGAGATACA	GAAATGAATT	GAGTGACATG	AGCAGGATGT	GGGGCCACCT	350
GAGCGAGGGG	TATGGCCAGC	TGTGCAGCAT	CTACCTGAAA	CTGCTAAGAA	400
CCAAGATGGA	GTACCACACC	AAAAATCCCA	GGTCCCCAGG	CAACCTGCAG	450
ATGAGTGACC	GCCAGCTGGA	CGAGGCTGGA	GAAAGTGACG	TGAACAACCT	500
TTTCCAGTTA	ACAGTGGAGA	TGTTTGACTA	CCTGGAGTGT	GAACCTCAACC	550
TCTTCCAAAC	AGTATTCAAC	TCCCTGGACA	TGTCCCGCTC	TGTGTCCGTG	600
ACGGCAGCAG	GGCAGTGCCG	CCTCGCCCCG	CTGATCCAGG	TCATCTTGGA	650
CTGCAGCCAC	CTTTATGACT	ACACTGTCAA	GCTTCTCTTC	AAACTCCACT	700
CCTGCCTCCC	AGCTGACACC	CTGCAAGGCC	ACCGGGACCG	CTTCATGGAG	750

CAGTTTACAA	AGTTGAAAGA	TCTGTTCTAC	CGCTCCAGCA	ACCTGCAGTA	800
CTTCAAGCGG	CTCATTCAGA	TCCCCCAGCT	GCCTGAGAAC	CCACCCAACCT	850
TCCTGCGAGC	CTCAGCCCTG	TCAGAACATA	TCAGCCCTGT	GGTGGTGATC	900
CCTGCAGAGG	CCTCATCCCC	CGACAGCGAG	CCAGTCCTAG	AGAAGGATGA	950
CCTCATGGAC	ATGGATGCCT	CTCAGCAGAA	TTTATTTGAC	AACAAGTTTG	1000
ATGACATCTT	TGGCAGTTCA	TTCAGCAGTG	ATCCCTTCAA	TTTCAACAGT	1050
CAAAATGGTG	TGAACAAGGA	TGAGAAGGAC	CACTTAATTG	AGCGACTATA	1100
CAGAGAGATC	AGTGGATTGA	AGGCACAGCT	AGAAAAATG	AAGACTGAGA	1150
GCCAGCGGGT	TGTGCTGCAG	CTGAAGGGCC	ACGTCAGCGA	GCTGGAAGCA	1200
GATCTGGCCG	AGCAGCAGCA	CCTGCGGCAG	CAGGCGGCCG	ACGACTGTGA	1250
ATTCCTGCGG	GCAGAACTGG	ACGAGCTCAG	GAGGCAGCGG	GAGGACACCG	1300
AGAAGGCTCA	GCGGAGCCTG	TCTGAGATAG	AAAGGAAAGC	TCAAGCCAAT	1350
GAACAGCGAT	ATAGCAAGCT	AAAGGAGAAG	TACAGCGAGC	TGGTTCAGAA	1400
CCACGCTGAC	CTGCTGCGGA	AGAATGCAGA	GGTGACCAA	CAGGTGTCCA	1450
TGGCCAGACA	AGCCAGGTA	GATTTGGAAC	GAGAGAAAAA	AGAGCTGGAG	1500
GATTTCGTTG	AGCGCATCAG	TGACCAGGGC	CAGCGGAAGA	CTCAAGAACA	1550
GCTGGAAGTT	CTAGAGAGCT	TGAAGCAGGA	ACTTGGCACA	AGCCAACGGG	1600
AGCTTCAGGT	TCTGCAAGGC	AGCCTGGAAG	CTTCTGCCCA	GTCAGAAGCA	1650
AACTGGGCAG	CCGAGTTCGC	CGAGCTAGAG	AAGGAGCGGG	ACAGCCTGGT	1700
GAGTGGCGCA	GCTCATAGGG	AGGAGGAATT	ATCTGCTCTT	CGGAAAGAAC	1750
TGCAGGACAC	TCAGCTCAAA	CTGGCCAGCA	CAGAGGAATC	TATGTGCCAG	1800
CTTGCCAAAG	ACCAACGAAA	AATGCTTCTG	GTGGGGTCCA	GGAAGGCTGC	1850
GGAGCAGGTG	ATACAAGACG	CCCTGAACCA	GCTTGAAGAA	CCTCCTCTCA	1900
TCAGCTGCGC	TGGGTCTGCA	GATCACCTCC	TCTCCACGGT	CACATCCATT	1950
TCCAGCTGCA	TCGAGCAACT	GGAGAAAAGC	TGGAGCCAGT	ATCTGGCCTG	2000
CCCAGAAGAC	ATCAGTGAGC	TTCTCCATTC	CATAACCCTG	CTGGCCCACT	2050
TGACCAGCGA	CGCCATTGCT	CATGGTGCCA	CCACCTGCCT	CAGAGCCCCA	2100
CCTGAGCCTG	CCGACTCACT	GACCGAGGCC	TGTAAGCAGT	ATGGCAGGGA	2150
AACCCTCGCC	TACCTGGCCT	CCCTGGAGGA	AGAGGGAAGC	CTTGAGAATG	2200
CCGACAGCAC	AGCCATGAGG	AACTGCCTGA	GCAAGATCAA	GGCCATCGGC	2250
GAGGAGCTCC	TGCCCAGGGG	ACTGGACATC	AAGCAGGAGG	AGCTGGGGGA	2300
CCTGGTGGAC	AAGGAGATGG	CGGCCACTTC	AGCTGCTATT	GAAACTTGCA	2350
CGGCCAGAAT	AGAGGAGATG	CTCAGCAAAT	CCCGAGCAGG	AGACACAGGA	2400
GTCAAATTGG	AGGTGAATGA	AAGGATCCTT	CGTTGCTGTA	CCAGCCTCAT	2450
GCAAGCTATT	CAGGTGCTCA	TCGTGGCCTC	TAAGGACCTC	CAGAGAGAGA	2500
TTGTGGAGAG	CGGCAGGGGT	ACAGCATCCC	CTAAAGAGTT	TTATGCCAAG	2550
AACTCTCGAT	GGACAGAAGG	ACTTATCTCA	GCCTCCAAGG	CTGTGGGCTG	2600
GGGAGCCACT	GTCATGGTGG	ATGCAGCTGA	TCTGGTGGTA	CAAGGCAGAG	2650
GGAAATTTGA	GGAGCTAATG	GTGTGTTCTC	ATGAAATTGC	TGCTAGCACA	2700
GCCCAGCTTG	TGGCTGCATC	CAAGGTGAAA	GCTGATAAGG	ACAGCCCCAA	2750
CCTAGCCCAG	CTGCAGCAGG	CCTCTCGGGG	AGTGAACCAG	GCCACTGCCG	2800
GCGTTGTGGC	CTCAACCATT	TCCGGCAAAT	CACAGATCGA	AGAGACAGAC	2850
AACATGGACT	TCTCAAGCAT	GACGCTGACA	CAGATCAAAC	GCCAAGAGAT	2900
GGATTCTCAG	GTTAGGGTGC	TAGAGCTAGA	AAATGAATTG	CAGAAGGAGC	2950
GTCAAAAAC	GGGAGAGCTT	CGGAAAAAGC	ACTACGAGCT	TGCTGGTGT	3000
GCTGAGGGCT	GGGAAGAAGG	AACAGAGGCA	TCTCCACCTA	CACTGCAAGA	3050
AGTGGTAACC	GAAAAAGAAT	AGAGCCAAAC	CAACACCCCA	TATGTCAGTG	3100
TAAATCCTTG	TTACCTATCT	CGTGTGTGTT	ATTTCCCCAG	CCACAGGCCA	3150
AATCCTTGGA	GTCCCAGGGG	CAGCCACACC	ACTGCCATTA	CCCAGTGCCG	3200
AGGACATGCA	TGACACTTCC	CAAAGATCCC	TCCATAGCGA	CACCCTTTCT	3250
GTTTGGACCC	ATGGTCATCT	CTGTTCTTTT	CCCGCCTCCC	TAGTTAGCAT	3300

CCAGGCTGGC	CAGTGCTGCC	CATGAGCAAG	CCTAGGTACG	AAGAGGGGTG	3350
GTGGGGGGCA	GGGCCACTCA	ACAGAGAGGA	CCAACATCCA	GTCCTGCTGA	3400
CTATTTGACC	CCCACAACAA	TGGGTATCCT	TAATAGAGGA	GCTGCTTGTT	3450
GTTTGTGAC	AGCTTGGA	GGGAAGATCT	TATGCCTTTT	CTTTTCTGTT	3500
TTCTTCTCAG	TCTTTTCAGT	TTCATCATTT	GCACAAACTT	GTGAGCATCA	3550
GAGGGCTGAT	GGATTCCAAA	CCAGGACACT	ACCCTGAGAT	CTGCACAGTC	3600
AGAAGGACGG	CAGGAGTGTC	CTGGCTGTGA	ATGCCAAAGC	CATTCTCCCC	3650
CTCTTTGGGC	AGTGCCATGG	ATTTCCACTG	CTTCTTATGG	TGGTTGGTTG	3700
GGTTTTTTGG	TTTTGTFTTT	TTTTTTTAAG	TTTCACTCAC	ATAGCCAACT	3750
CTCCCAAAGG	GCACACCCCT	GGGGCTGAGT	CTCCAGGGCC	CCCCAACTGT	3800
GGTAGCTCCA	GCGATGGTGC	TGCCCAGGCC	TCTCGGTGCT	CCATCTCCGC	3850
CTCCCACTG	ACCAAGTGCT	GGCCCACCCA	GTCCATGCTC	CAGGGTCAGG	3900
CGGAGCTGCT	GAGTGACAGC	TTTCCTCAAA	AAGCAGAAGG	AGAGTGAGTG	3950
CCTTTCCTC	CTAAAGCTGA	ATCCCGCGG	AAAGCCTCTG	TCCGCCTTTA	4000
CAAGGGAGAA	GACAACAGAA	AGAGGGACAA	GAGGGTTCAC	ACAGCCCAGT	4050
TCCCGTGACG	AGGCTCAAAA	ACTTGATCAC	ATGCTTGAAT	GGAGCTGGTG	4100
AGATCAACAA	CACTACTTCC	CTGCCGGAAT	GAAGTGTCCG	TGAATGGTCT	4150
CTGTCAAGCG	GGCCGTCTCC	CTTGCCCCAG	AGACGGAGTG	TGGGAGTGAT	4200
TCCCAACTCC	TTTCTGCAGA	CGTCTGCCTT	GGCATCCTCT	TGAATAGGAA	4250
GATCGTTCCA	CTTTCTACGC	AATTGACAAA	CCCGGAAGAT	CAGATGCAAT	4300
TGCTCCCATC	AGGGAAGAAC	CCTATACTTG	GTTTGCTACC	CTTAGTATTT	4350
ATTACTAACC	TCCCTTAAGC	AGCAACAGCC	TACAAAGAGA	TGCTTGGAGC	4400
AATCAGAACT	TCAGGTGTGA	CTCTAGCAAA	GCTCATCTTT	CTGCCCCGCT	4450
ACATCAGCCT	TCAAGAATCA	GAAGAAAGCC	AAGGTGCTGG	ACTGTTACTG	4500
ACTTGGATCC	CAAAGCAAGG	AGATCATTTG	GAGCTCTTGG	GTCAGAGAAA	4550
ATGAGAAAGG	ACAGAGCCAG	CGGCTCCAAC	TCCTTTCAGC	CACATGCCCC	4600
AGGCTCTCGC	TGCCCTGTGG	ACAGGATGAG	GACAGAGGGC	ACATGAACAG	4650
CTTGCCAGGG	ATGGGCAGCC	CAACAGCACT	TTTCCTCTTC	TAGATGGACC	4700
CCAGCATTTA	AGTGACCTTC	TGATCTTGGG	AAAACAGCGT	CTTCCTTCTT	4750
TATCTATAGC	AACTCATTGG	TGGTAGCCAT	CAAGCACTTC	GGAATT	4796

(2) INFORMATION FOR SEQ ID NO: 4:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 924

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:

Met	Ser	Arg	Met	Trp	Gly	His	Leu	Ser	Glu	Gly	Tyr	Gly	Gln	Leu
1				5					10					15

Cys	Ser	Ile	Tyr	Leu	Lys	Leu	Leu	Arg	Thr	Lys	Met	Glu	Tyr	His
				20					25					30

Thr	Lys	Asn	Pro	Arg	Phe	Pro	Gly	Asn	Leu	Gln	Met	Ser	Asp	Arg	
				35					40					45	
Gln	Leu	Asp	Glu	Ala	Gly	Glu	Ser	Asp	Val	Asn	Asn	Phe	Phe	Gln	
				50					55					60	
Leu	Thr	Val	Glu	Met	Phe	Asp	Tyr	Leu	Glu	Cys	Glu	Leu	Asn	Leu	
				65					70					75	
Phe	Gln	Thr	Val	Phe	Asn	Ser	Leu	Asp	Met	Ser	Arg	Ser	Val	Ser	
				80					85					90	
Val	Thr	Ala	Ala	Gly	Gln	Cys	Arg	Leu	Ala	Pro	Leu	Ile	Gln	Val	
				95					100					105	
Ile	Leu	Asp	Cys	Ser	His	Leu	Tyr	Asp	Tyr	Thr	Val	Lys	Leu	Leu	
				110					115					120	
Phe	Lys	Leu	His	Ser	Cys	Leu	Pro	Ala	Asp	Thr	Leu	Gln	Gly	His	
				125					130					135	
Arg	Asp	Arg	Phe	Met	Glu	Gln	Phe	Thr	Lys	Leu	Lys	Asp	Leu	Phe	
				140					145					150	
Tyr	Arg	Ser	Ser	Asn	Leu	Gln	Tyr	Phe	Lys	Arg	Leu	Ile	Gln	Ile	
				155					160					165	
Pro	Gln	Leu	Pro	Glu	Asn	Pro	Pro	Asn	Phe	Leu	Arg	Ala	Ser	Ala	
				170					175					180	
Leu	Ser	Glu	His	Ile	Ser	Pro	Val	Val	Val	Ile	Pro	Ala	Glu	Ala	
				185					190					195	
Ser	Ser	Pro	Asp	Ser	Glu	Pro	Val	Leu	Glu	Lys	Asp	Asp	Leu	Met	
				200					205					210	
Asp	Met	Asp	Ala	Ser	Gln	Gln	Asn	Leu	Phe	Asp	Asn	Lys	Phe	Asp	
				215					220					225	
Asp	Ile	Phe	Gly	Ser	Ser	Phe	Ser	Ser	Asp	Pro	Phe	Asn	Phe	Asn	
				230					235					240	
Ser	Gln	Asn	Gly	Val	Asn	Lys	Asp	Glu	Lys	Asp	His	Leu	Ile	Glu	
				245					250					255	
Arg	Leu	Tyr	Arg	Glu	Ile	Ser	Gly	Leu	Lys	Ala	Gln	Leu	Glu	Asn	
				260					265					270	
Met	Lys	Thr	Glu	Ser	Gln	Arg	Val	Val	Leu	Gln	Leu	Lys	Gly	His	
				275					280					285	

Val	Ser	Glu	Leu	Glu	Ala	Asp	Leu	Ala	Glu	Gln	Gln	His	Leu	Arg
				290					295					300
Gln	Gln	Ala	Ala	Asp	Asp	Cys	Glu	Phe	Leu	Arg	Ala	Glu	Leu	Asp
				305					310					315
Glu	Leu	Arg	Arg	Gln	Arg	Glu	Asp	Thr	Glu	Lys	Ala	Gln	Arg	Ser
				320					325					330
Leu	Ser	Glu	Ile	Glu	Arg	Lys	Ala	Gln	Ala	Asn	Glu	Gln	Arg	Tyr
				335					340					345
Ser	Lys	Leu	Lys	Glu	Lys	Tyr	Ser	Glu	Leu	Val	Gln	Asn	His	Ala
				350					355					360
Asp	Leu	Leu	Arg	Lys	Asn	Ala	Glu	Val	Thr	Lys	Gln	Val	Ser	Met
				365					370					375
Ala	Arg	Gln	Ala	Gln	Val	Asp	Leu	Glu	Arg	Glu	Lys	Lys	Glu	Leu
				380					385					390
Glu	Asp	Ser	Leu	Glu	Arg	Ile	Ser	Asp	Gln	Gly	Gln	Arg	Lys	Thr
				395					400					405
Gln	Glu	Gln	Leu	Glu	Val	Leu	Glu	Ser	Leu	Lys	Gln	Glu	Leu	Gly
				410					415					420
Thr	Ser	Gln	Arg	Glu	Leu	Gln	Val	Leu	Gln	Gly	Ser	Leu	Glu	Thr
				425					430					435
Ser	Ala	Gln	Ser	Glu	Ala	Asn	Trp	Ala	Ala	Glu	Phe	Ala	Glu	Leu
				440					445					450
Glu	Lys	Glu	Arg	Asp	Ser	Leu	Val	Ser	Gly	Ala	Ala	His	Arg	Glu
				455					460					465
Glu	Glu	Leu	Ser	Ala	Leu	Arg	Lys	Glu	Leu	Gln	Asp	Thr	Gln	Leu
				470					475					480
Lys	Leu	Ala	Ser	Thr	Glu	Glu	Ser	Met	Cys	Gln	Leu	Ala	Lys	Asp
				485					490					495
Gln	Arg	Lys	Met	Leu	Leu	Val	Gly	Ser	Arg	Lys	Ala	Ala	Glu	Gln
				500					505					510
Val	Ile	Gln	Asp	Ala	Leu	Asn	Gln	Leu	Glu	Glu	Pro	Pro	Leu	Ile
				515					520					525
Ser	Cys	Ala	Gly	Ser	Ala	Asp	His	Leu	Leu	Ser	Thr	Val	Thr	Ser
				530					535					540

Ile	Ser	Ser	Cys	Ile	Glu	Gln	Leu	Glu	Lys	Ser	Trp	Ser	Gln	Tyr
				545					550					555
Leu	Ala	Cys	Pro	Glu	Asp	Ile	Ser	Gly	Leu	Leu	His	Ser	Ile	Thr
				560					565					570
Leu	Leu	Ala	His	Leu	Thr	Ser	Asp	Ala	Ile	Ala	His	Gly	Ala	Thr
				575					580					585
Thr	Cys	Leu	Arg	Ala	Pro	Pro	Glu	Pro	Ala	Asp	Ser	Leu	Thr	Glu
				590					595					600
Ala	Cys	Lys	Gln	Tyr	Gly	Arg	Glu	Thr	Leu	Ala	Tyr	Leu	Ala	Ser
				605					610					615
Leu	Glu	Glu	Glu	Gly	Ser	Leu	Glu	Asn	Ala	Asp	Ser	Thr	Ala	Met
				620					625					630
Arg	Asn	Cys	Leu	Ser	Lys	Ile	Lys	Ala	Ile	Gly	Glu	Glu	Leu	Leu
				635					640					645
Pro	Arg	Gly	Leu	Asp	Ile	Lys	Gln	Glu	Glu	Leu	Gly	Asp	Leu	Val
				650					655					660
Asp	Lys	Glu	Met	Ala	Ala	Thr	Ser	Ala	Ala	Ile	Glu	Thr	Cys	Thr
				665					670					675
Ala	Arg	Ile	Glu	Glu	Met	Leu	Ser	Lys	Ser	Arg	Ala	Gly	Asp	Thr
				680					685					690
Gly	Val	Lys	Leu	Glu	Val	Asn	Glu	Arg	Ile	Leu	Arg	Cys	Cys	Thr
				695					700					705
Ser	Leu	Met	Gln	Ala	Ile	Gln	Val	Leu	Ile	Val	Ala	Ser	Lys	Asp
				710					715					720
Leu	Gln	Arg	Glu	Ile	Val	Glu	Ser	Gly	Arg	Gly	Thr	Ala	Ser	Pro
				725					730					735
Lys	Glu	Phe	Tyr	Ala	Lys	Asn	Ser	Arg	Trp	Thr	Glu	Gly	Leu	Ile
				740					745					750
Ser	Ala	Ser	Lys	Ala	Val	Gly	Trp	Gly	Ala	Thr	Val	Met	Val	Asp
				765					770					775
Ala	Ala	Asp	Leu	Val	Val	Gln	Gly	Arg	Gly	Lys	Phe	Glu	Glu	Leu
				780					785					790
Met	Val	Cys	Ser	His	Glu	Ile	Ala	Ala	Ser	Thr	Ala	Gln	Leu	Val
				795					800					805

Ala	Ala	Ser	Lys	Val	Lys	Ala	Asp	Lys	Asp	Ser	Pro	Asn	Leu	Ala
				810					815					820
Gln	Leu	Gln	Gln	Ala	Ser	Arg	Gly	Val	Asn	Gln	Ala	Thr	Ala	Gly
				825					830					835
Val	Val	Ala	Ser	Thr	Ile	Ser	Gly	Lys	Ser	Gln	Ile	Glu	Glu	Thr
				840					845					850
Asp	Asn	Met	Asp	Phe	Ser	Ser	Met	Thr	Leu	Thr	Gln	Ile	Lys	Arg
				855					860					865
Gln	Glu	Met	Asp	Ser	Gln	Val	Arg	Val	Leu	Glu	Leu	Glu	Asn	Glu
				870					875					880
Leu	Gln	Lys	Glu	Arg	Gln	Lys	Leu	Gly	Glu	Leu	Arg	Lys	Lys	His
				885					890					895
Tyr	Glu	Leu	Ala	Gly	Val	Ala	Glu	Gly	Trp	Glu	Glu	Gly	Thr	Glu
				900					905					910
Ala	Ser	Pro	Pro	Thr	Leu	Gln	Glu	Val	Val	Thr	Glu	Lys	Glu	
				915					920				924	

(2) INFORMATION FOR SEQ ID NO: 5

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1090

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:

Met	Leu	Leu	Cys	Gln	Gly	Ser	Glu	Trp	Arg	Arg	Asp	Gln	Gln	Leu
				5					10					15
Gly	Thr	Ala	Asn	Ala	Arg	Gln	Trp	Cys	Pro	Leu	Pro	Gln	Asp	Ala
				20					25					30
Gln	Pro	Ala	Gly	Ser	Trp	Glu	Arg	Cys	Pro	Pro	Leu	Pro	Pro	Ala
				35					40					45
Gly	Arg	Leu	Gln	Gly	Thr	Asp	His	Pro	Trp	Gly	Trp	Gly	Arg	Leu
				50					55					60

Ala	Gly	Gly	Gly	Glu	Arg	Gly	Gly	Leu	Trp	Glu	Gly	Leu	Ser	His
				65					70					75
Ser	Gln	Arg	Leu	Ile	His	Leu	Ile	Leu	Leu	Ser	Leu	Pro	Leu	Leu
				80					85					90
Val	Phe	Gln	Thr	Val	Ser	Ile	Asn	Lys	Ala	Ile	Asn	Thr	Gln	Glu
				95					100					105
Val	Ala	Val	Lys	Glu	Lys	His	Ala	Arg	Thr	Cys	Ile	Leu	Gly	Thr
				110					115					120
His	His	Glu	Lys	Gly	Ala	Gln	Thr	Phe	Trp	Ser	Val	Val	Asn	Arg
				125					130					135
Leu	Pro	Leu	Ser	Ser	Asn	Ala	Val	Leu	Cys	Trp	Lys	Phe	Cys	His
				140					145					150
Val	Phe	His	Lys	Leu	Leu	Arg	Asp	Gly	His	Pro	Asn	Val	Leu	Lys
				155					160					165
Asp	Ser	Leu	Arg	Tyr	Arg	Asn	Glu	Leu	Ser	Asp	Met	Ser	Arg	Met
				170					175					180
Trp	Gly	His	Leu	Ser	Glu	Gly	Tyr	Gly	Gln	Leu	Cys	Ser	Ile	Tyr
				185					190					195
Leu	Lys	Leu	Leu	Arg	Thr	Lys	Met	Glu	Tyr	His	Thr	Lys	Asn	Pro
				200					205					210
Arg	Phe	Pro	Gly	Asn	Leu	Gln	Met	Ser	Asp	Arg	Gln	Leu	Asp	Glu
				215					220					225
Ala	Gly	Glu	Ser	Asp	Val	Asn	Asn	Phe	Phe	Gln	Leu	Thr	Val	Glu
				230					235					240
Met	Phe	Asp	Tyr	Leu	Glu	Cys	Glu	Leu	Asn	Leu	Phe	Gln	Thr	Val
				245					250					255
Phe	Asn	Ser	Leu	Asp	Met	Ser	Arg	Ser	Val	Ser	Val	Thr	Ala	Ala
				260					265					270
Gly	Gln	Cys	Arg	Leu	Ala	Pro	Leu	Ile	Gln	Val	Ile	Leu	Asp	Cys
				275					288					285
Ser	His	Leu	Tyr	Asp	Tyr	Thr	Val	Lys	Leu	Leu	Phe	Lys	Leu	His
				290					295					300
Ser	Cys	Leu	Pro	Ala	Asp	Thr	Leu	Gln	Gly	His	Arg	Asp	Arg	Phe
				305					310					315

Met	Glu	Gln	Phe	Thr	Lys	Leu	Lys	Asp	Leu	Phe	Tyr	Arg	Ser	Ser	
				320					325					330	
Asn	Leu	Gln	Tyr	Phe	Lys	Arg	Leu	Ile	Gln	Ile	Pro	Gln	Leu	Pro	
				335					340					345	
Glu	Asn	Pro	Pro	Asn	Phe	Leu	Arg	Ala	Ser	Ala	Leu	Ser	Glu	His	
				350					355					360	
Ile	Ser	Pro	Val	Val	Val	Ile	Pro	Ala	Glu	Ala	Ser	Ser	Pro	Asp	
				365					370					375	
Ser	Glu	Pro	Val	Leu	Glu	Lys	Asp	Asp	Leu	Met	Asp	Met	Asp	Ala	
				380					385					390	
Ser	Gln	Gln	Asn	Leu	Phe	Asp	Asn	Lys	Phe	Asp	Asp	Ile	Phe	Gly	
				395					400					405	
Ser	Ser	Phe	Ser	Ser	Asp	Pro	Phe	Asn	Phe	Asn	Ser	Gln	Asn	Gly	
				410					415					420	
Val	Asn	Lys	Asp	Glu	Lys	Asp	His	Leu	Ile	Glu	Arg	Leu	Tyr	Arg	
				425					430					435	
Glu	Ile	Ser	Gly	Leu	Lys	Ala	Gln	Leu	Glu	Asn	Met	Lys	Thr	Glu	
				440					445					450	
Ser	Gln	Arg	Val	Val	Leu	Gln	Leu	Lys	Gly	His	Val	Ser	Glu	Leu	
				455					460					465	
Glu	Ala	Asp	Leu	Ala	Glu	Gln	Gln	His	Leu	Arg	Gln	Gln	Ala	Ala	
				470					475					480	
Asp	Asp	Cys	Glu	Phe	Leu	Arg	Ala	Glu	Leu	Asp	Glu	Leu	Arg	Arg	
				485					490					495	
Gln	Arg	Glu	Asp	Thr	Glu	Lys	Ala	Gln	Arg	Ser	Leu	Ser	Glu	Ile	
				500					505					510	
Glu	Arg	Lys	Ala	Gln	Ala	Asn	Glu	Gln	Arg	Tyr	Ser	Lys	Leu	Lys	
				515					520					525	
Glu	Lys	Tyr	Ser	Glu	Leu	Val	Gln	Asn	His	Ala	Asp	Leu	Leu	Arg	
				530					535					540	
Lys	Asn	Ala	Glu	Val	Thr	Lys	Gln	Val	Ser	Met	Ala	Arg	Gln	Ala	
				545					550					555	
Gln	Val	Asp	Leu	Glu	Arg	Glu	Lys	Lys	Glu	Leu	Glu	Asp	Ser	Leu	
				560					565					570	

Glu	Arg	Ile	Ser	Asp	Gln	Gly	Gln	Arg	Lys	Thr	Gln	Glu	Gln	Leu
				575					588					585
Glu	Val	Leu	Glu	Ser	Leu	Lys	Gln	Glu	Leu	Ala	Thr	Ser	Gln	Arg
				590					595					600
Glu	Leu	Gln	Val	Leu	Gln	Gly	Ser	Leu	Glu	Thr	Ser	Ala	Gln	Ser
				605					610					615
Glu	Ala	Asn	Trp	Ala	Ala	Glu	Phe	Ala	Glu	Leu	Glu	Lys	Glu	Arg
				620					625					630
Asp	Ser	Leu	Val	Ser	Gly	Ala	Ala	His	Arg	Glu	Glu	Glu	Leu	Ser
				635					640					645
Ala	Leu	Arg	Lys	Glu	Leu	Gln	Asp	Thr	Gln	Leu	Lys	Leu	Ala	Ser
				650					655					660
Thr	Glu	Glu	Ser	Met	Cys	Gln	Leu	Ala	Lys	Asp	Gln	Arg	Lys	Met
				665					670					675
Leu	Leu	Val	Gly	Ser	Arg	Lys	Ala	Ala	Glu	Gln	Val	Ile	Gln	Asp
				680					685					690
Ala	Leu	Asn	Gln	Leu	Glu	Glu	Pro	Pro	Leu	Ile	Ser	Cys	Ala	Gly
				695					700					705
Ser	Ala	Asp	His	Leu	Leu	Ser	Thr	Val	Thr	Ser	Ile	Ser	Ser	Cys
				710					715					720
Ile	Glu	Gln	Leu	Glu	Lys	Ser	Trp	Ser	Gln	Tyr	Leu	Ala	Cys	Pro
				725					730					735
Glu	Asp	Ile	Ser	Gly	Leu	Leu	His	Ser	Ile	Thr	Leu	Leu	Ala	His
				740					745					750
Leu	Thr	Ser	Asp	Ala	Ile	Ala	His	Gly	Ala	Thr	Thr	Cys	Leu	Arg
				755					760					765
Ala	Pro	Pro	Glu	Pro	Ala	Asp	Ser	Leu	Thr	Glu	Ala	Cys	Lys	Gln
				770					775					780
Tyr	Gly	Arg	Glu	Thr	Leu	Ala	Tyr	Leu	Ala	Ser	Leu	Glu	Glu	Glu
				785					790					795
Gly	Ser	Leu	Glu	Asn	Ala	Asp	Ser	Thr	Ala	Met	Arg	Asn	Cys	Leu
				800					805					810
Ser	Lys	Ile	Lys	Ala	Ile	Gly	Glu	Glu	Leu	Leu	Pro	Arg	Gly	Leu
				815					820					825

Asp	Ile	Lys	Gln	Glu	Glu	Leu	Gly	Asp	Leu	Val	Asp	Lys	Glu	Met
				830					835					840
Ala	Ala	Thr	Ser	Ala	Ala	Ile	Glu	Thr	Ala	Thr	Ala	Arg	Ile	Glu
				845					850					855
Glu	Met	Leu	Ser	Lys	Ser	Arg	Ala	Gly	Asp	Thr	Gly	Val	Lys	Leu
				860					865					870
Glu	Val	Asn	Glu	Arg	Ile	Leu	Gly	Cys	Cys	Thr	Ser	Leu	Met	Gln
				875					888					885
Ala	Ile	Gln	Val	Leu	Ile	Val	Ala	Ser	Lys	Asp	Leu	Gln	Arg	Glu
				890					895					900
Ile	Val	Glu	Ser	Gly	Arg	Gly	Thr	Ala	Ser	Pro	Lys	Glu	Phe	Tyr
				905					910					915
Ala	Lys	Asn	Ser	Arg	Trp	Thr	Glu	Gly	Leu	Ile	Ser	Ala	Ser	Lys
				920					925					930
Ala	Val	Gly	Trp	Gly	Ala	Thr	Val	Met	Val	Asp	Ala	Ala	Asp	Leu
				935					940					945
Val	Val	Gln	Gly	Arg	Gly	Lys	Phe	Glu	Glu	Leu	Met	Val	Cys	Ser
				950					955					960
His	Glu	Ile	Ala	Ala	Ser	Thr	Ala	Gln	Leu	Val	Ala	Ala	Ser	Lys
				965					970					975
Val	Lys	Ala	Asp	Lys	Asp	Ser	Pro	Asn	Leu	Ala	Gln	Leu	Gln	Gln
				980					985					990
Ala	Ser	Arg	Gly	Val	Asn	Gln	Ala	Thr	Ala	Gly	Val	Val	Ala	Ser
				995					1000					1005
Thr	Ile	Ser	Gly	Lys	Ser	Gln	Ile	Glu	Glu	Thr	Asp	Asn	Met	Asp
				1010					1015					1020
Phe	Ser	Ser	Met	Thr	Leu	Thr	Gln	Ile	Lys	Arg	Gln	Glu	Met	Asp
				1025					1030					1035
Ser	Gln	Val	Arg	Val	Leu	Glu	Leu	Glu	Asn	Glu	Leu	Gln	Lys	Glu
				1040					1045					1050
Arg	Gln	Lys	Leu	Gly	Glu	Leu	Arg	Lys	Lys	His	Tyr	Glu	Leu	Ala
				1055					1060					1065
Gly	Val	Ala	Glu	Gly	Trp	Glu	Glu	Gly	Thr	Glu	Ala	Ser	Pro	Pro
				1070					1075					1080

Thr Leu Gln Glu Val Val Thr Glu Lys Glu
1085 1090

(2) INFORMATION FOR SEQ ID NO:6:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 3301

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: cDNA for Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:

CGGTGAGCTG	GAGGAGCAGC	GGAAGCAGAA	GCAGAAGGCC	CTGGTGGATA	50
ATGAGCAGCT	CCGCCACGAG	CTGGCCCAGC	TGAGGGCTGC	CCAGCTGGAG	100
CGCGAGCGGA	GCCAGGGCCT	GCGTGAGGAG	GCTGAGAGGA	AGGCCAGTGC	150
CACGGAGGCG	CGCTACAACA	AGCTGAAGGA	AAAGCACAGT	GAGCTCGTCC	200
ATGTGCACGC	GGAGCTGCTC	AGAAAGAACG	CGGACACAGC	CAAGCAGCTG	250
ACGGTGACGC	AGCAAAGCCA	GGAGGAGGTG	GCGCGGGTGA	AGGAGCAGCT	300
GGCCTTCCAG	GTGGAGCAGG	TGAAGCGGGA	GTCGGAGTTG	AAGCTAGAGG	350
AGAAGAGCGA	CCAGCAGGAG	AAGCTCAAGA	GGGAGCTGGA	GGCCAAGGCC	400
GGAGAGCTGG	CCCGCGCGCA	GGAGGCCCTG	AGCCACACAG	AGCAGAGCAA	450
GTCGGAGCTG	AGCTCACGGC	TGGACACACT	GAGTGCGGAG	AAGGATGCTC	500
TGAGTGAGGC	TGTGCGGCAG	CGGGAGGCAG	ACCTGCTGGC	GGCGCAGAGC	550
CTGGTGCGCG	AGACAGAGGC	GGCGCTGAGC	CGGGAGCAGC	AGCGCAGCTC	600
CCAGGAGCAG	GGCGAGTTGC	AGGGCCCGCT	GGCAGAGAGG	GAGTCTCAGG	650
AGCAGGGGCT	GCGGCAGAGG	CTGCTGGACG	AGCAGTTCGC	AGTGTTGCGG	700
GGCGCTGCTG	CCGAGGCCGC	GGGCATCCTG	CAGGATGCCG	TGAGCAAGCT	750
GGACGACCCC	CTGCACCTGC	GCTGTACCAG	CTCCCCAGAC	TACCTGGTGA	800
GCAGGGCCCA	GGAGGCCCTG	GATGCCGTGA	GCACCCTGGA	GGAGGGCCAC	850
GCCCAGTACC	TGACCTCCTT	GGCAGACGCC	TCCGCCCTGG	TGGCAGCTCT	900
GACCCGCTTC	TCCCACCTGG	CTGCGGATAC	CATCATCAAT	GGCGGTGCCA	950
CCTCGCACCT	GGCTCCCACC	GACCCTGCCG	ACCGCCTCAT	AGACACCTGC	1000
AGGGAGTGCG	GGGCCCCGGC	TCTGGAGCTC	ATGGGGCAGC	TGCAGGACCA	1050
GCAGGCTCTG	CGGCACATGC	AGGCCAGCCT	GGTGCGGACA	CCCCTGCAGG	1100
GCATCCTTCA	GCTGGGCCAA	GAAGTGAAC	CCAAGAGCCT	AGATGTGCGG	1150
CAGGAGGAGC	TGGGGGCCGT	GGTCGACAAG	GAGATGGCGG	CCACATCCGC	1200
AGCCATTGAA	GATGCTGTGC	GGAGGATTGA	GGACATGATG	AACCAGGCAC	1250
GCCACGCCAG	CTCGGGGGTG	AAGCTGGAGG	TGAACGAGAG	GATCCTCAAC	1300
TCCTGCACAG	ACCTGATGAA	GGCTATCCGG	CTCCTGGTGA	CGACATCCAC	1350
TAGCCTGCAG	AAGGAGATCG	TGGAGAGCGG	CAGGGGGGCA	GCCACGCAGC	1400
AGGAATTTTA	CGCCAAGAAC	TCGCGCTGGA	CCGAAGGCCT	CATCTCGGCC	1450
TCCAAGGCTG	TGGGCTGGGG	AGCCACACAG	CTGGTGGAGG	CAGCTGACAA	1500
GGTGGTGCTT	CACACGGGCA	AGTATGAGGA	GCTCATCGTC	TGCTCCCACG	1550
AGATCGCAGC	CAGCACGGCC	CAGCTGGTGG	CGGCCCTCAA	GGTGAAGGCC	1600

AACAAGCACA	GCCCCCACCT	GAGCCGCTGT	CAGGAATGTT	CTCGCACAGT	1650
CAATGAGAGG	GCTGCCAATG	TGGTGGCCTC	CACCAAGTCA	GGCCAGGAGC	1700
AGATTGAGGA	CAGAGACACC	ATGGATTCT	CCGGCCTGTC	CCTCATCAAG	1750
CTGAAGAAGC	AGGAGATGGA	GACGCAGGTG	CGTGTCTCTG	AGCTGGAGAA	1800
GACGCTGGAG	GCTGAACGCA	TGCGGCTGGG	GGAGTTGCGG	AAGCAACACT	1850
ACGTGCTGGC	TGGGGCATCA	GGCAGCCCTG	GAGAGGAGGT	GGCCATCCGG	1900
CCCAGCACTG	CCCCCCGAAG	TGTAACCACC	AAGAAACCAC	CCCTGGCCCA	1950
GAAGCCCAGC	GTGGCCCCCA	GACAGGACCA	CCAGCTTGAC	AAAAAGGATG	2000
GCATCTACCC	AGCTCAACTC	GTGAACTACT	AGGCCCCCCA	GGGGTCCAGC	2050
AGGGTGGCTG	GTGACAGGCC	TGGGCCTCTG	CAACTGCCCT	GACAGGACCG	2100
AGAGGCCTTG	CCCCTCCACC	TGGTGCCCAA	GCCTCCCGCC	CCACCGTCTG	2150
GATCAATGTC	CTCAAGGCC	CTGGCCCTTA	CTGAGCCTGC	AGGGTCCTGG	2200
GCCATGTGGG	TGGTGCTTCT	GGATGTGAGT	CTCTTATTTA	TCTGCAGAAG	2250
GAACTTTGGG	GTGCAGCCAG	GACCCGGTAG	GCCTGAGCCT	CAACTCTTCA	2300
GAAAATAGTG	TTTTTAATAT	TCCTCTTCAG	AAAATAGTGT	TTTTAATATT	2350
CCGAGCTAGA	GCTCTTCTTC	CTACGTTTGT	AGTCAGCACA	CTGGGAAACC	2400
GGGCCAGCGT	GGGGCTCCCT	GCCTTCTGGA	CTCCTGAAGG	TCGTGGATGG	2450
ATGGAAGGCA	CACAGCCCGT	GCCGGCTGAT	GGGACGAGGG	TCAGGCATCC	2500
TGTCTGTGGC	CTTCTGGGGC	ACCGATTCTA	CCAGGCCCTC	CAGCTGCGTG	2550
GTCTCCGCAG	ACCAGGCTCT	GTGTGGGCTA	GAGGAATGTC	GCCCATTACC	2600
TCCTCAGGCC	CTGGCCCTCG	GGCCTCCGTG	ATGGGAGCCC	CCCAGGAGGG	2700
GTCAGATGCT	GGAAGGGGCC	GCTTTCTGGG	GAGTGAGGTG	AGACATAGCG	2750
GCCCAGGCGC	TGCCTTCACT	CCTGGAGTTT	CCATTTCAG	CTGGAATCTG	2800
CAGCCACCCC	CATTTCCCTGT	TTTCCATTCC	CCCGTTCTGG	CCGCGCCCCA	2850
CTGCCCACCT	GAAGGGGTGG	TTTCCAGCCC	TCCGGAGAGT	GGGCTTGGCC	2900
CTAGGCCCTC	CAGCTCAGCC	AGAAAAAGCC	CAGAAACCCA	GGTGCTGGAC	2950
CAGGGCCCTC	AGGGAGGGAC	CCTGCGGCTA	GAGTGGGCTA	GGCCCTGGCT	3000
TTGCCCCGTC	GATTTGAACG	AATGTGTGTC	CCTTGAGCCC	AAGGAGAGCG	3050
GCAGGAGGGG	TGGGACCAGG	CTGGGAGGAC	AGAGCCAGCA	GCTGCCATGC	3100
CCTCCTGCTC	CCCCACCCC	AGCCCTAGCC	CTTTAGCCTT	TCACCCTGTG	3150
CTCTGGAAAG	GCTACCAAAT	ACTGGCCAAG	GTCAGGAGGA	GCAAAAATGA	3200
GCCAGCACCA	GCGCCTTGGC	TTTGTGTTAG	CATTTCCCTC	TGAAGTGTTT	3250
TGTTGGCAAT	AAAATGCACT	TTGACTGTTA	AAAAAAAAAA	AAAAAAAAAA	3300
A					3301

(2) INFORMATION FOR SEQ ID NO: 7

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 676

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:

Gly Glu Leu Glu Glu Gln Arg Lys Gln Lys Glu Lys Ala Leu Val

Asp	Asn	Glu	Gln	Leu	Arg	His	Glu	Leu	Ala	Gln	Leu	Arg	Ala	Ala	
				20					25					30	
Gln	Leu	Glu	Arg	Glu	Arg	Ser	Gln	Gly	Leu	Arg	Glu	Glu	Ala	Glu	
				35					40					45	
Arg	Lys	Ala	Ser	Ala	Thr	Glu	Ala	Arg	Tyr	Asn	Lys	Leu	Lys	Glu	
				50					55					60	
Lys	His	Ser	Glu	Leu	Val	His	Val	His	Ala	Glu	Leu	Leu	Arg	Lys	
				65					70					75	
Asn	Ala	Asp	Thr	Ala	Lys	Gln	Leu	Thr	Val	Thr	Gln	Gln	Ser	Gln	
				80					85					90	
Glu	Glu	Val	Ala	Arg	Val	Lys	Glu	Gln	Leu	Ala	Phe	Gln	Val	Glu	
				95					100					105	
Gln	Val	Lys	Arg	Glu	Ser	Glu	Leu	Lys	Leu	Glu	Glu	Lys	Ser	Asp	
				110					115					120	
Gln	Gln	Glu	Lys	Leu	Lys	Arg	Glu	Leu	Glu	Ala	Lys	Ala	Gly	Glu	
				125					130					135	
Leu	Ala	Arg	Ala	Gln	Glu	Ala	Leu	Ser	His	Thr	Glu	Gln	Ser	Lys	
				140					145					150	
Ser	Glu	Leu	Ser	Ser	Arg	Leu	Asp	Thr	Leu	Ser	Ala	Glu	Lys	Asp	
				155					160					165	
Ala	Leu	Ser	Gly	Ala	Val	Arg	Gln	Arg	Glu	Ala	Asp	Leu	Leu	Ala	
				170					175					180	
Ala	Gln	Ser	Leu	Val	Arg	Glu	Thr	Glu	Ala	Ala	Leu	Ser	Arg	Glu	
				185					190					195	
Gln	Gln	Arg	Ser	Ser	Gln	Glu	Gln	Gly	Glu	Leu	Gln	Gly	Arg	Leu	
				200					205					210	
Ala	Glu	Arg	Glu	Ser	Gln	Glu	Gln	Gly	Leu	Arg	Gln	Arg	Leu	Leu	
				215					220					225	
Asp	Glu	Gln	Phe	Ala	Val	Leu	Arg	Gly	Ala	Ala	Ala	Glu	Ala	Ala	
				230					235					240	
Gly	Ile	Leu	Gln	Asp	Ala	Val	Ser	Lys	Leu	Asp	Asp	Pro	Leu	His	
				245					250					255	
Leu	Arg	Cys	Thr	Ser	Ser	Pro	Asp	Tyr	Leu	Val	Ser	Arg	Ala	Gln	
				260					265					270	

Glu	Ala	Leu	Asp	Ala	Val	Ser	Thr	Leu	Glu	Glu	Gly	His	Ala	Gln
				275					288					285
Tyr	Leu	Thr	Ser	Leu	Ala	Asp	Ala	Ser	Ala	Leu	Val	Ala	Ala	Leu
				290					295					300
Thr	Arg	Phe	Ser	His	Leu	Ala	Ala	Asp	Thr	Ile	Ile	Asn	Gly	Gly
				305					310					315
Ala	Thr	Ser	His	Leu	Ala	Pro	Thr	Asp	Pro	Ala	Asp	Arg	Leu	Ile
				320					325					330
Asp	Thr	Cys	Arg	Glu	Cys	Gly	Ala	Arg	Ala	Leu	Glu	Leu	Met	Gly
				335					340					345
Gln	Leu	Gln	Asp	Gln	Gln	Ala	Leu	Arg	His	Met	Gln	Ala	Ser	Leu
				350					355					360
Val	Arg	Thr	Pro	Leu	Gln	Gly	Ile	Leu	Gln	Leu	Gly	Gln	Glu	Leu
				365					370					375
Lys	Pro	Lys	Ser	Leu	Asp	Val	Arg	Gln	Glu	Glu	Leu	Gly	Ala	Val
				380					385					390
Val	Asp	Lys	Glu	Met	Ala	Ala	Thr	Ser	Ala	Ala	Ile	Glu	Asp	Ala
				395					400					405
Val	Arg	Arg	Ile	Glu	Asp	Met	Met	Asn	Gln	Ala	Arg	His	Ala	Ser
				410					415					420
Ser	Gly	Val	Lys	Leu	Glu	Val	Asn	Glu	Arg	Ile	Leu	Asn	Ser	Cys
				425					430					435
Thr	Asp	Leu	Met	Lys	Ala	Ile	Arg	Leu	Leu	Val	Thr	Thr	Ser	Thr
				440					445					450
Ser	Leu	Gln	Lys	Glu	Ile	Val	Glu	Ser	Gly	Arg	Gly	Ala	Ala	Thr
				455					460					465
Gln	Gln	Glu	Phe	Tyr	Ala	Lys	Asn	Ser	Arg	Trp	Thr	Glu	Gly	Leu
				470					475					480
Ile	Ser	Ala	Ser	Lys	Ala	Val	Gly	Trp	Gly	Ala	Thr	Gln	Leu	Val
				485					490					495
Glu	Ala	Ala	Asp	Lys	Val	Val	Leu	His	Thr	Gly	Lys	Tyr	Glu	Glu
				500					505					510
Leu	Ile	Val	Cys	Ser	His	Glu	Ile	Ala	Ala	Ser	Thr	Ala	Gln	Leu
				515					520					525

Val	Ala	Ala	Ser	Lys	Val	Lys	Ala	Asn	Lys	His	Ser	Pro	His	Leu
				530					535					540
Ser	Arg	Leu	Gln	Glu	Cys	Ser	Arg	Thr	Val	Asn	Glu	Arg	Ala	Ala
				545					550					555
Asn	Val	Val	Ala	Ser	Thr	Lys	Ser	Gly	Gln	Glu	Gln	Ile	Glu	Asp
				560					565					570
Arg	Asp	Thr	Met	Asp	Phe	Ser	Gly	Leu	Ser	Leu	Ile	Lys	Leu	Lys
				575					588					585
Lys	Gln	Glu	Met	Glu	Thr	Gln	Val	Arg	Val	Leu	Glu	Leu	Glu	Lys
				590					595					600
Thr	Leu	Glu	Ala	Glu	Arg	Met	Arg	Leu	Gly	Glu	Leu	Arg	Lys	Gln
				605					610					615
His	Tyr	Val	Leu	Ala	Gly	Ala	Ser	Gly	Ser	Pro	Gly	Glu	Glu	Val
				620					625					630
Ala	Ile	Arg	Pro	Ser	Thr	Ala	Pro	Arg	Ser	Val	Thr	Thr	Lys	Lys
				635					640					645
Pro	Pro	Leu	Ala	Gln	Lys	Pro	Ser	Val	Ala	Pro	Arg	Gln	Asp	His
				650					655					660
Gln	Leu	Asp	Lys	Lys	Asp	Gly	Ile	Tyr	Pro	Ala	Gln	Leu	Val	Asn
				665					670					675

Tyr

(2) INFORMATION FOR SEQ ID NO:8:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 2338
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA
- (iii) HYPOTHETICAL: no
- (iv) ANTI-SENSE: no
- (vi) ORIGINAL SOURCE:

- (A) ORGANISM: mouse
- (ix) FEATURE: cDNA for Huntingtin-interacting protein - mHIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:

GGCACGAGGG	CTCATTCAGA	TCCCCCAGCT	GCCCCGAGAAT	CCACCCAACTT	50
CCTACGAGCC	TCGGCCCTGT	CAGAGCACAT	CAGTCCTGTG	GTGGTGATCCC	100
GGCAGAGGTG	TCATCCCCAG	ACAGTGAGCC	TGTCCTGGAG	AAGGATGACCT	150
CATGGACATG	GACGCCTCCC	AGCAGACTTT	GTTTGACAAC	AAGTTTGATGA	200

CGTCTTTGGC	AGCTCATTGA	GCAGCGACCC	TTTCAATTTC	AACAATCAAAA	250
TGGCGTGAAC	AAGGACGAGA	AGGACCACTT	GATTGAACGC	CTGTACAGAGA	300
GATCAGTGGA	CTGACAGGGC	AGCTGGACAA	CATGAAGATT	GAGAGCCAGCG	350
GGCCATGCTG	CAGCTGAAGG	GTCGAGTGAG	TGAGCTGGAG	GCAGAGCTAGC	400
AGAGCAGCAG	CACTTGGGCC	GGCAGGCTAT	GGATGACTGC	GAGTTCCTGCG	450
CACTGAGCTG	GATGAACTGA	AGAGGCAGCG	AGAGGACACG	GAGAAGGCACA	500
GCGCAGCCTG	ACTGAGATAG	AAAGAAAGGC	CCAGGCTAAT	GAACAGAGGTA	550
TAGCAAGTTA	AAAGAGAAGT	ACAGTGAACT	GGTGCAGAAC	CATGCTGACCT	600
GCTGCGGAAG	AACGCAGAGG	TGACCAAACA	GGTGTCCGTG	GCCCGGCAAGC	650
CCAGGTGGAT	TTGGAAAGAG	AGAAAAAAGA	GCTAGCAGAT	TCCTTTGCAC	700
GTGTAAGTGA	CCAGGCCAG	CGGAAGACTC	AAGAGCAACA	GGATGTTCTA	750
GAGAACCTGA	AGCATGAACT	GGCCACCAGC	AGACAGGAGC	TGCAGGTCCT	800
CCACAGCAAC	CTGGAAACCT	CTGCCCAGTC	AGAAGCGAAA	TGGCTGACAC	850
AGATCGCCGA	GTTGGAGAAG	GAACAAGGCA	GCTTGGCGAC	TGTTGCAGCT	900
CAGAGAGAGG	AAGAGTTATC	AGCCCTCCGA	GACCAGCTGG	AAAGCACCCA	950
GATCAAGCTG	GCTGGGGCCC	AGGAATCCAT	GTGCCAGCAG	GTGAAGGACC	1000
AGAGGAAAAC	CCTCTTGGA	GGGATCAGGA	AGGCTGCGGA	GCGTGAGATA	1050
CAGGAGGCGC	TGAGCCAGCT	TGAGGAACCC	ACCCTCATCA	GCTGTGCAGG	1100
ATCCACAGAT	CACCTTCTCT	CCAAAGTCAG	CTCCGTTTCC	AGCTGCCTCG	1150
AGCAACTGGA	AAAGAACGGC	AGCCAGTATC	TGGCCTGCCC	AGAAGATATT	1200
AGTGAGCTTC	TGCACTCGAT	CACCCTGCTT	GCCCCTTGA	CCGGTGACAC	1250
TGTCATCCAG	GGGAGTGCCA	CCAGCCTCCG	GGCCCCACCG	GAGCCAGCCG	1300
ACTCGTTGAC	GGAGGCCTGT	AGGCAGTATG	GCAGAGAAAC	CCTGGCCTAT	1350
CTGTCCTCCC	TGGAGGAAGA	GGGAAGTGTG	GAGAATGCTG	ACGTCACAGC	1400
CCTTAGGAAT	TGCCTCAGCA	GGGTCAAGAC	CCTTGGCGAG	GAGCTGCTGC	1450
CCAGGGGCCT	GGACATCAAG	CAGGAAGAGC	TGGGTGACCT	GGTGGACAAG	1500
GAGATGGCAG	CCACTTCAGC	TGCCATTGAA	GCTGCCACCA	CCCGGATAGA	1550
GGAAATTCTC	AGTAAGTCCC	GAGCAGGAGA	CACGGGAGTC	AAGCTGGAGG	1600
TGAATGAGAG	GATCCTGGGT	TCCTGTACCA	GCCTGATGCA	GGCCATCAAG	1650
GTGCTCGTTG	TGGCCTCCAA	GGACCTCCAG	AAGGAGATAG	TGGAGAGTGG	1700
CAGGGGTAGT	GCATCCCCTA	AAGAATTTTA	CGCCAAGAAC	TCTCGGTGGA	1750
CGGAAGGGCT	GATATCCGCC	TCCAAAGCTG	TTGGTTGGGG	AGCTACCATC	1800
ATGGTGATG	CTGCTGATCT	TGTGGTCCAA	GGCAAAGGGA	AGTTCGAGGA	1850
GCTGATGGTG	TGTTACGCG	AGATTGCTGC	CAGTACTGCC	CAGCTCGTGG	1900
CTGCATCCAA	GGTGAAAGCG	AACAAGGGCA	GCCTCAATCT	GACCCAGCTG	2000
CAGCAGGCCT	CTCGAGGAGT	GAACCAGGCC	ACAGCCGCTG	TGGTGGCCTC	2050
AACCATTTCT	GGCAAATCTC	AGATTGAGGA	AACAGACAGT	ATGGACTTCT	2100
CAAGCATGAC	ACTGACCCAG	ATCAAGCGCC	AGGAGATGGA	TTCCCAGGTT	2150
AGGGTGCTGG	AGCTGGAAAA	TGACCTGCAG	AAGGAGCGTC	AGAAACTAGG	2200
AGAGCTACGG	AAGAAACACT	ACGAGCTGGA	GGGCGTGGCT	GAGGGCTGGG	2250
AGGAAGGGAC	AGAAGCATCA	CCGTCTACTG	TCCAAGAAGC	AATACCGGAC	2300
AAAGAGTAGA	GCCAAGCCGA	CACCCACAC	ATCAGAAA		2338

(2) INFORMATION FOR SEQ ID NO: 9:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 676

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: mouse

(ix) FEATURE: Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9:

Ala	Arg	Gly	Leu	Ile	Gln	Ile	Pro	Gln	Leu	Pro	Glu	Asn	Pro	Pro	5	10	15
Asn	Phe	Leu	Arg	Ala	Ser	Ala	Leu	Ser	Glu	His	Ile	Ser	Pro	Val	20	25	30
Val	Val	Ile	Pro	Ala	Glu	Val	Ser	Ser	Pro	Asp	Ser	Glu	Pro	Val	35	40	45
Leu	Glu	Lys	Asp	Asp	Leu	Met	Asp	Met	Asp	Ala	Ser	Gln	Gln	Thr	50	55	60
Leu	Phe	Asp	Asn	Lys	Phe	Asp	Asp	Val	Phe	Gly	Ser	Ser	Leu	Ser	65	70	75
Ser	Asp	Pro	Phe	Asn	Phe	Asn	Asn	Gln	Asn	Gly	Val	Asn	Lys	Asp	80	85	90
Glu	Lys	Asp	His	Leu	Ile	Glu	Arg	Leu	Tyr	Arg	Glu	Ile	Ser	Gly	95	100	105
Leu	Thr	Gly	Gln	Leu	Asp	Asn	Met	Lys	Ile	Glu	Ser	Gln	Arg	Ala	110	115	120
Met	Leu	Gln	Leu	Lys	Gly	Arg	Val	Ser	Glu	Leu	Glu	Ala	Glu	Leu	125	130	135
Ala	Glu	Gln	Gln	His	Leu	Gly	Arg	Gln	Ala	Met	Asp	Asp	Cys	Glu	140	145	150
Phe	Leu	Arg	Thr	Glu	Leu	Asp	Glu	Leu	Lys	Arg	Gln	Arg	Glu	Asp	155	160	165
Thr	Glu	Lys	Ala	Gln	Arg	Ser	Leu	Thr	Glu	Ile	Glu	Arg	Lys	Ala	170	175	180
Gln	Ala	Asn	Glu	Gln	Arg	Tyr	Ser	Lys	Leu	Lys	Glu	Lys	Tyr	Ser	185	190	195
Glu	Leu	Val	Gln	Asn	His	Ala	Asp	Leu	Leu	Arg	Lys	Asn	Ala	Glu	200	205	210
Val	Thr	Lys	Gln	Val	Ser	Val	Ala	Arg	Gln	Ala	Gln	Val	Asp	Leu	215	220	225
Glu	Arg	Glu	Lys	Lys	Glu	Leu	Ala	Asp	Ser	Phe	Ala	Arg	Val	Ser			

	230		235		240
Asp Gln Ala Gln Arg Lys Thr Gln Glu	Gln Gln Asp Val Leu Glu				
245	250				255
Asn Leu Lys His Glu Leu Ala Thr Ser	Arg Gln Glu Leu Gln Val				
260	265				270
Leu His Ser Asn Leu Glu Thr Ser Ala	Gln Ser Glu Ala Lys Trp				
275	288				285
Leu Thr Gln Ile Ala Glu Leu Glu Lys	Glu Gln Gly Ser Leu Ala				
290	295				300
Thr Val Ala Ala Gln Arg Glu Glu Glu	Leu Ser Ala Leu Arg Asp				
305	310				315
Gln Leu Glu Ser Thr Gln Ile Lys Leu	Ala Gly Ala Gln Glu Ser				
320	325				330
Met Cys Gln Gln Val Lys Asp Gln Arg	Lys Thr Leu Leu Ala Gly				
335	340				345
Ile Arg Lys Ala Ala Glu Arg Glu Ile	Gln Glu Ala Leu Ser Gln				
350	355				360
Leu Glu Glu Pro Thr Leu Ile Ser Cys	Ala Gly Ser Thr Asp His				
365	370				375
Leu Leu Ser Lys Val Ser Ser Val Ser	Ser Cys Leu Glu Gln Leu				
380	385				390
Glu Lys Asn Gly Ser Gln Tyr Leu Ala	Cys Pro Glu Asp Ile Ser				
395	400				405
Glu Leu Leu His Ser Ile Thr Leu Leu	Ala His Leu Thr Gly Asp				
410	415				420
Thr Val Ile Gln Gly Ser Ala Thr Ser	Leu Arg Ala Pro Pro Glu				
425	430				435
Pro Ala Asp Ser Leu Thr Glu Ala Cys	Arg Gln Tyr Gly Arg Glu				
440	445				450
Thr Leu Ala Tyr Leu Ser Ser Leu Glu	Glu Glu Gly Thr Val Glu				
455	460				465
Asn Ala Asp Val Thr Ala Leu Arg Asn	Cys Leu Ser Arg Val Lys				
470	475				480

Thr	Leu	Gly	Glu	Glu	Leu	Leu	Pro	Arg	Gly	Leu	Asp	Ile	Lys	Gln
				485					490					495
Glu	Glu	Leu	Gly	Asp	Leu	Val	Asp	Lys	Glu	Met	Ala	Ala	Thr	Ser
				500					505					510
Ala	Ala	Ile	Glu	Ala	Ala	Thr	Thr	Arg	Ile	Glu	Glu	Ile	Leu	Ser
				515					520					525
Lys	Ser	Arg	Ala	Gly	Asp	Thr	Gly	Val	Lys	Leu	Glu	Val	Asn	Glu
				530					535					540
Arg	Ile	Leu	Gly	Ser	Cys	Thr	Ser	Leu	Met	Gln	Ala	Ile	Lys	Val
				545					550					555
Leu	Val	Val	Ala	Ser	Lys	Asp	Leu	Gln	Lys	Glu	Ile	Val	Glu	Ser
				560					565					570
Gly	Arg	Gly	Ser	Ala	Ser	Pro	Lys	Glu	Phe	Tyr	Ala	Lys	Asn	Ser
				575					588					585
Arg	Trp	Thr	Glu	Gly	Leu	Ile	Ser	Ala	Ser	Lys	Ala	Val	Gly	Trp
				590					595					600
Gly	Ala	Thr	Ile	Met	Val	Asp	Ala	Ala	Asp	Leu	Val	Val	Gln	Gly
				605					610					615
Lys	Gly	Lys	Phe	Glu	Glu	Leu	Met	Val	Cys	Ser	Arg	Glu	Ile	Ala
				620					625					630
Ala	Ser	Thr	Ala	Gln	Leu	Val	Ala	Ala	Ser	Lys	Val	Lys	Ala	Asn
				635					640					645
Lys	Gly	Ser	Leu	Asn	Leu	Thr	Gln	Leu	Gln	Gln	Ala	Ser	Arg	Gly
				650					655					660
Val	Asn	Gln	Ala	Thr	Ala	Ala	Val	Val	Ala	Ser	Thr	Ile	Ser	Gly
				665					670					675
Lys	Ser	Gln	Ile	Glu	Glu	Thr	Asp	Ser	Met	Asp	Phe	Ser	Ser	Met
				680					685					690
Thr	Leu	Thr	Gln	Ile	Lys	Arg	Gln	Glu	Met	Asp	Ser	Gln	Val	Arg
				695					700					705
Val	Leu	Glu	Leu	Glu	Asn	Asp	Leu	Gln	Lys	Glu	Arg	Gln	Lys	Leu
				710					715					720
Gly	Glu	Leu	Arg	Lys	Lys	His	Tyr	Glu	Leu	Glu	Gly	Val	Ala	Glu
				725					730					735

Gly Trp Glu Glu Gly Thr Glu Ala Ser Pro Ser Thr Val Gln Glu
740 745 750

Ala Ile Pro Asp Lys Glu
755

(2) INFORMATION FOR SEQ ID NO:10:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 3964

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: mouse

(ix) FEATURE: cDNA for Huntingtin-interacting protein - mHIP1a

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:

GGCACGAGGC	GGCGCGCGGC	CTCCGTGTGC	CTAGGCTTGA	GGCGGGCGGT	50
GACGCCTCAT	TCGCGCGGAG	CCGGGCCGGG	ACACGGTCGG	CGGCAGCATG	100
AACAGCATCA	AGAATGTGCC	GGCGCGGGTG	CTGAGCCGCA	GGCCGGGCCA	150
CAGCCTAGAG	GCCGAGCGCG	AGCAGTTCGA	CAAGACGCAG	GCCATCAGTA	200
TCAGCAAAGC	CATCAACAGC	CAGGAGGCC	CAGTGAAGGA	GAAGCATGCC	250
CGGCGTATCA	TCCTGGGCAC	GCATCATGAG	AAGGGAGCCT	TCACCTTCTG	300
GTCCTATGCC	ATCGGCCTGC	CGCTGTCCAG	CAGCTCCATC	CTCAGCTGGA	350
AGTTCTGTCA	CGTCCTTCAC	AAGTCCCTCC	GGGACGGACA	CCCCAACGTC	400
CTGCATGACT	ATCAGCGGTA	CCGAGCAAC	ATACGTGAGA	TCGGTGACTT	450
GTGGGGCCAC	CTTCGTGACC	AGTATGGACA	CCTGGTGAAT	ATCTATACCA	500
AACTGTTGCT	GACTAAGATC	TCCTTCCACC	TTAAGCACCC	CCAGTTTCTT	550
GCAGGCCTGG	AGGTAACAGA	TGAGGTGTTG	GAGAAGGCGG	CGGGAAGTGA	600
TGTCAACAAC	ATTTTTCAGC	TTACCGTGGA	GATGTTTGAC	TACATGGACT	650
GTGAACTGAA	GCTTTCTGAG	TCAGTTTTCC	GGCAGCTCAA	CACGGCCATC	700
GCAGTGTCCC	AGATGTCTTC	TGGCCAGTGT	CGCCTAGCGC	CGCTCATCCA	750
GGTCATTGAG	GACTGCAGCC	ACCTGTACCA	CTACACAGTG	AAGCTCATGT	800
TTAAGCTGCA	CTCCTGTCTC	CCGGCAGACA	CCCTGCAAGG	CCACAGGGAT	850
CGGTTCACAG	AGCAGTTCCA	CAGCCTCAAA	AACTTCTTCC	GCCGGGCTTC	900
AGACATGCTG	TACTTCAAGA	GGCTCATCCA	GATCCCAGCG	CTGCCTGAGG	950
GACCCCCCAA	TTTCCTGCGG	GCTTCAGCCC	TGGCTGAGCA	CATCAAGCCG	1000
GTGGTGGTGA	TTCCCGAGGA	GGCCCCAGAG	GAAGAGGAGC	CTGAGAACCCT	1050
AATTGAAATC	AGCAGTGCGC	CCCCTGCTGG	GGAGCCAGTG	GTGGTGGCTG	1100
ACCTCTTTGA	TCAGACCTTT	GGACCCCCCA	ATGGCTCCAT	GAAGGATGAC	1150
AGGGACCTCC	AAATCGAGAA	CTTGAAGAGA	GAGGTGGAGA	CCCTCCGTGC	1200
TGAGCTGGAG	AAGATTAAGA	TGGAGGCACA	GCGGTACATC	TCCCAGCTGA	1250
AGGGCCAGGT	GAATGGCCTG	GAGGCAGAGC	TGGAGGAGCA	GCGCAAGCAG	1300
AAGCAGAAGG	CCCTGGTGGA	CAACGAGCAG	CTGCGCCACG	AGCTGGCCCA	1350
GCTCAAGGCC	CTGCAGCTGG	AGGGCGCCCG	CAACCAGGGC	CTTCGAGAGG	1400
AAGCAGAGAG	GAAGGCCAGT	GCCACGGAGG	CACGCTACAG	CAAGCTGAAG	1450
GAGAAACACA	GCGAACTCAT	TAACACGCAC	GCCGAGCTGC	TCAGGAAGAA	1500

CGCAGACACG	GCCAAGCAGC	TGACAGTGAC	ACAGCAGAGC	CAGGAGGAGG	1550
TGGCACGGGT	AAAGGAACAG	CTGGCCTTCC	AGATGGAGCA	AGCGAAGCGT	1600
GAGTCTGAGA	TGAAGATGGA	AGAGCAGAGC	GACCAGTTGG	AGAAGCTCAA	1650
GAGGGAGCTG	GCGGCCAGGG	CAGGAGAGCT	GGCCCGTGCG	CAGGAGGCCC	1700
TGAGCCGCAC	AGAACAGAGT	GGGTCAGAGC	TGAGCTCACG	GCTGGACACA	1750
CTGAACGCGG	AGAAGGAAGC	CCTGAGTGGA	GTCGTTGCGC	AGCGTGAGGC	1800
AGAGCTGCTG	GCCGCTCAGA	GCCTGGTGCG	GGAGAAGGAG	GAGGCGCTTA	1850
GCCAAGAGCA	GCAGCGGAGC	TCCCAGGAGA	AGGGCGAGCT	ACGGGGGCGC	1900
CTGGCAGAAA	AGGAGTCTCA	GGAGCAGGGG	CTTCGGCAGA	AGCTGCTGGA	1950
TGAGCAGTTG	GCGGTGTTGC	GAAGTGCAGC	CGCCGAGGCA	GAGGCCATCC	2000
TACAGGATGC	AGTGAGCAAG	CTGGACGACC	CCCTGCACCT	CCGCTGCACC	2050
AGCTCCCCAG	ACTACTTGGT	GAGCCGGGCT	CAGGCAGCCC	TGGACAGCGT	2100
GAGCGGCCTG	GAGCAGGGCC	ACACCCAGTA	CCTGGCTTCC	TCCGAAGATG	2150
CTTCTGCCCT	GGTGGCAGCG	CTGACCCGCT	TCTCCCATT	GGCTGCGGAC	2200
ACCATTGTCA	ATGGTGCCGC	CACCTCCCAC	CTGGCCCCCA	CCGACCCCGC	2250
CGACCGCCTG	ATGGACACAT	GCAGGGAGTG	TGGAGCCCGG	GCTCTGGAGC	2300
TGGTGGGACA	GCTGCAAGAC	CAGACAGTGC	TACGGAGGGC	TCAGCCCAGC	2350
CTGATGCGGG	CCCCCTGCA	GGGCATTCTG	CAGTTGGGCC	AGGACTTGAA	2400
GCCTAAGAGC	CTGGATGTAC	GGCAAGAGGA	GCTAGGGGCC	ATGGTGGACA	2450
AGGAGATGGC	GGCCACCTCG	GCAGCCATTG	AGGACGCTGT	GCGGAGGATC	2500
GAGGACATGA	TGAGCCAGGC	CCGCCACGAG	AGCTCAGGCG	TGAAACTGGA	2550
GGTGAATGAG	AGGATCCTCA	ACTCCTGCAC	AGACCTGATG	AAGGCTATCC	2600
GGCTCCTGGT	GATGACCTCC	ACCAGCCTGC	AGAAGGAAAT	TGTGGAGAGC	2650
GGCAGGGGGG	CAGCAACGCA	GCAGGAATTT	TATGCCAAGA	ATTCACGGTG	2700
GACTGAAGGC	CTCATCTCAG	CCTCTAAGGC	AGTGGGCTGG	GGAGCCACAC	2750
AGCTGGTGGA	GTCAGCTGAC	AAGGTTGTGC	TTCACATGGG	CAAATACGAG	2800
GAACTCATCG	TCTGCTCCCA	TGAGATTGCG	GCCAGCACGG	CCCAGCTGGT	2850
GGCAGCCTCG	AAGGTGAAAG	CCAACAAGAA	CAGTCCCCAC	TTGAGCCGCC	2900
TGCAGGAATG	TTCCCGCACT	GTCAACGAGA	GGGCTGCCAA	CGTCGTGGCC	2950
TCCACCAAAT	CTGGCCAGGA	GCAGATTGAG	GACAGAGACA	CCATGGATTT	3000
CTCTGGCCTG	TCCCTCATCA	AGTTGAAGAA	GCAGGAGATG	GAGACACAGG	3050
TGCGAGTCTT	GGAGCTGGAG	AAGACACTAG	AGGCAGAGCG	TGTCCGGCTC	3100
GGGGAGCTTC	GGAAACAGCA	CTATGTACTG	GCTGGGGGGA	TGGGAACACC	3150
TAGCGAAGAA	GAACCCAGCA	GACCCAGCCC	AGCTCCCCGA	AGTGGGGCCA	3200
CTAAGAAGCC	ACCGCTGGCC	CAGAAACCCA	GCATAGCCCC	CAGGACAGAC	3250
AACCAGCTCGA	CAAAAAGGAT	GGTGTCTACC	CAGCTCAACT	TGTGAAGTAC	3300
TAGGCCCCCTAA	GGTGTTCAGC	AGGATGGCTG	GTGGTTGTGC	CTGGGCTTCA	3350
TGTGGCTGTCT	GGCAGTGGTC	AAGGGGCCTC	TGAGAAGCCT	CCAACTCCTG	3400
CCCAAGGGGCC	TAGTCTGTGG	GACAGTTCAT	CTGGATGTGA	ATCTATTTAT	3450
CTTAAGTAGGA	ACTGCCTCGA	GCAGCTGGGA	CCCAGCAGGC	CTGAGCCACA	3500
AATCTGCAGCG	GACATCAGAG	ATAGTCTGAA	TGCTGCGAGG	TATTTCTTTC	3550
TTCGTAAGTTT	AGTCAGCACA	CTGGGAAAAG	GTCACATAAG	CCAGGAGCCT	3600
CCTTGTCTCTG	GACTCAAAAG	TCTGAGGCCT	TAAGTGAACA	ACAGAAAGAG	3650
GGTCCCTGCTG	GCTACCAGGG	ATAAGGGGAT	GACCTGTGAC	CCTTGAGCCA	3700
GGGAGAGCAGG	TAAGCTGGGT	GGTGTTCATCA	CCTGGGGGCC	TGGTGCTAGG	3750
GCATCCATGCT	GGGAGCCCCA	GGAGACCAGG	CTTTGTGTGG	GAGCCTGGCA	3800
TCATCGTGGCT	GGGGCAGCCC	CTGCTCAGGT	GCTGTCTCTG	CCCGTGACCT	3850
TGAAGCCACCC	TCCCCCGTA	CAGTTTTCCA	TTCTCCTGGC	TACTAGTGTG	3900
GCTGTTCATTG	CCTACCTTGA	TGAGTAGATT	TCAGCCCTCC	TAAAGCTGGG	3950
GCCTTTCCTCG	TGCC				3964

(2) INFORMATION FOR SEQ ID NO: 11:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 676

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: mouse

(ix) FEATURE: Huntingtin-interacting protein -mHIP1a

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:11:

Met	Asn	Ser	Ile	Lys	Asn	Val	Pro	Ala	Arg	Val	Leu	Ser	Arg	Arg	
				5					10					15	
Pro	Gly	His	Ser	Leu	Glu	Ala	Glu	Arg	Glu	Gln	Phe	Asp	Lys	Thr	
				20					25					30	
Gln	Ala	Ile	Ser	Ile	Ser	Lys	Ala	Ile	Asn	Ser	Gln	Glu	Ala	Pro	
				35					40					45	
Val	Lys	Glu	Lys	His	Ala	Arg	Arg	Ile	Ile	Leu	Gly	Thr	His	His	
				50					55					60	
Glu	Lys	Gly	Ala	Phe	Thr	Phe	Trp	Ser	Tyr	Ala	Ile	Gly	Leu	Pro	
				65					70					75	
Leu	Ser	Ser	Ser	Ser	Ile	Leu	Ser	Trp	Lys	Phe	Cys	His	Val	Leu	
				80					85					90	
His	Lys	Val	Leu	Arg	Asp	Gly	His	Pro	Asn	Val	Leu	His	Asp	Tyr	
				95					100					105	
Gln	Arg	Tyr	Arg	Ser	Asn	Ile	Arg	Glu	Ile	Gly	Asp	Leu	Trp	Gly	
				110					115					120	
His	Leu	Arg	Asp	Gln	Tyr	Gly	His	Leu	Val	Asn	Ile	Tyr	Thr	Lys	
				125					130					135	
Leu	Leu	Leu	Thr	Lys	Ile	Ser	Phe	His	Leu	Lys	His	Pro	Gln	Phe	
				140					145					150	
Pro	Ala	Gly	Leu	Glu	Val	Thr	Asp	Glu	Val	Leu	Glu	Lys	Ala	Ala	
				155					160					165	
Gly	Thr	Asp	Val	Asn	Asn	Ile	Phe	Gln	Leu	Thr	Val	Glu	Met	Phe	
				170					175					180	
Asp	Tyr	Met	Asp	Cys	Glu	Leu	Lys	Leu	Ser	Glu	Ser	Val	Phe	Arg	

	185	190	195
Gln Leu Asn Thr	Ala Ile Ala Val Ser	Gln Met Ser Ser	Gly Gln
	200	205	210
Cys Arg Leu Ala	Pro Leu Ile Gln Val	Ile Gln Asp Cys	Ser His
	215	220	225
Leu Tyr His Tyr	Thr Val Lys Leu Met	Phe Lys Leu His	Ser Cys
	230	235	240
Leu Pro Ala Asp	Thr Leu Gln Gly His	Arg Asp Arg Phe	His Glu
	245	250	255
Gln Phe His Ser	Leu Lys Asn Phe Phe	Arg Arg Ala Ser	Asp Met
	260	265	270
Leu Tyr Phe Lys	Arg Leu Ile Gln Ile	Pro Arg Leu Pro	Glu Gly
	275	288	285
Pro Pro Asn Phe	Leu Arg Ala Ser Ala	Leu Ala Glu His	Ile Lys
	290	295	300
Pro Val Val Val	Ile Pro Glu Glu Ala	Pro Glu Glu Glu	Glu Pro
	305	310	315
Glu Asn Leu Ile	Glu Ile Ser Ser Ala	Pro Pro Ala Gly	Glu Pro
	320	325	330
Val Val Val Ala	Asp Leu Phe Asp Gln	Thr Phe Gly Pro	Pro Asn
	335	340	345
Gly Ser Met Lys	Asp Asp Arg Asp Leu	Gln Ile Glu Asn	Leu Lys
	350	355	360
Arg Glu Val Glu	Thr Leu Arg Ala Glu	Leu Glu Lys Ile	Lys Met
	365	370	375
Glu Ala Gln Arg	Tyr Ile Ser Gln Leu	Lys Gly Gln Val	Asn Gly
	380	385	390
Leu Glu Ala Glu	Leu Glu Glu Gln Arg	Lys Gln Lys Gln	Lys Ala
	395	400	405
Leu Val Asp Asn	Glu Gln Leu Arg His	Glu Leu Ala Gln	Leu Lys
	410	415	420
Ala Leu Gln Leu	Glu Gly Ala Arg Asn	Gln Gly Leu Arg	Glu Glu
	425	430	435
Ala Glu Arg Lys	Ala Ser Ala Thr Glu	Ala Arg Tyr Ser	Lys Leu

	440		445		450
Lys Glu Lys His	Ser Glu Leu Ile Asn	Thr His Ala Glu Leu Leu			
	455		460		465
Arg Lys Asn Ala	Asp Thr Ala Lys Gln	Leu Thr Val Thr Gln Gln			
	470		475		480
Ser Gln Glu Glu	Val Ala Arg Val Lys	Glu Gln Leu Ala Phe Gln			
	485		490		495
Met Glu Gln Ala	Lys Arg Glu Ser Glu	Met Lys Met Glu Glu Gln			
	500		505		510
Ser Asp Gln Leu	Glu Lys Leu Lys Arg	Glu Leu Ala Ala Arg Ala			
	515		520		525
Gly Glu Leu Ala	Arg Ala Gln Glu Ala	Leu Ser Arg Thr Glu Gln			
	530		535		540
Ser Gly Ser Glu	Leu Ser Ser Arg Leu	Asp Thr Leu Asn Ala Glu			
	545		550		555
Lys Glu Ala Leu	Ser Gly Val Val Arg	Gln Arg Glu Ala Glu Leu			
	560		565		570
Leu Ala Ala Gln	Ser Leu Val Arg Glu	Lys Glu Glu Ala Leu Ser			
	575		588		585
Gln Glu Gln Gln	Arg Ser Ser Gln Glu	Lys Gly Glu Leu Arg Gly			
	590		595		600
Gln Leu Ala Glu	Lys Glu Ser Gln Glu	Gln Gly Leu Arg Gln Lys			
	605		610		615
Leu Leu Asp Glu	Gln Leu Ala Val Leu	Arg Ser Ala Ala Ala Glu			
	620		625		630
Ala Glu Ala Ile	Leu Gln Asp Ala Val	Ser Lys Leu Asp Asp Pro			
	635		640		645
Leu His Leu Arg	Cys Thr Ser Ser Pro	Asp Tyr Leu Val Ser Arg			
	650		655		660
Ala Gln Ala Ala	Leu Asp Ser Val Ser	Gly Leu Glu Gln Gly His			
	665		670		675
Thr Gln Tyr Leu	Ala Ser Ser Glu Asp	Ala Ser Ala Leu Val Ala			
	680		685		690
Ala Leu Thr Arg	Phe Ser His Leu Ala	Ala Asp Thr Ile Val Asn			

695										700					705				
Gly	Ala	Ala	Thr	Ser	His	Leu	Ala	Pro	Thr	Asp	Pro	Ala	Asp	Arg					
				710					715					720					
Leu	Met	Asp	Thr	Cys	Arg	Glu	Cys	Gly	Ala	Arg	Ala	Leu	Glu	Leu					
				725					730					735					
Val	Gly	Gln	Leu	Gln	Asp	Gln	Thr	Val	Leu	Arg	Arg	Ala	Gln	Pro					
				740					745					750					
Ser	Leu	Met	Arg	Ala	Pro	Leu	Gln	Gly	Ile	Leu	Gln	Leu	Gly	Gln					
				755					760					765					
Asp	Leu	Lys	Pro	Lys	Ser	Leu	Asp	Val	Arg	Gln	Glu	Glu	Leu	Gly					
				770					775					780					
Ala	Met	Val	Asp	Lys	Glu	Met	Ala	Ala	Thr	Ser	Ala	Ala	Ile	Glu					
				785					790					795					
Asp	Ala	Val	Arg	Arg	Ile	Glu	Asp	Met	Met	Ser	Gln	Ala	Arg	His					
				800					805					810					
Glu	Ser	Ser	Gly	Val	Lys	Leu	Glu	Val	Asn	Glu	Arg	Ile	Leu	Asn					
				815					820					825					
Ser	Cys	Thr	Asp	Leu	Met	Lys	Ala	Ile	Arg	Leu	Leu	Val	Met	Thr					
				830					835					840					
Ser	Thr	Ser	Leu	Gln	Lys	Glu	Ile	Val	Glu	Ser	Gly	Arg	Gly	Ala					
				845					850					855					
Ala	Thr	Gln	Gln	Glu	Phe	Tyr	Ala	Lys	Asn	Ser	Arg	Trp	Thr	Glu					
				860					865					870					
Gly	Leu	Ile	Ser	Ala	Ser	Lys	Ala	Val	Gly	Trp	Gly	Ala	Thr	Gln					
				875					888					885					
Leu	Val	Glu	Ser	Ala	Asp	Lys	Val	Val	Leu	His	Met	Gly	Lys	Tyr					
				890					895					900					
Glu	Glu	Leu	Ile	Val	Cys	Ser	His	Glu	Ile	Ala	Ala	Ser	Thr	Ala					
				905					910					915					
Gln	Leu	Val	Ala	Ala	Ser	Lys	Val	Lys	Ala	Asn	Lys	Asn	Ser	Pro					
				920					925					930					
His	Leu	Ser	Arg	Leu	Gln	Glu	Cys	Ser	Arg	Thr	Val	Asn	Glu	Arg					
				935					940					945					
Ala	Ala	Asn	Val	Val	Ala	Ser	Thr	Lys	Ser	Gly	Gln	Glu	Gln	Ile					

	950	955	960
Glu Asp Arg Asp Thr Met Asp Phe Ser Gly Leu Ser Leu Ile Lys	965	970	975
Leu Lys Lys Gln Glu Met Glu Thr Gln Val Arg Val Leu Glu Leu	980	985	990
Glu Lys Thr Leu Glu Ala Glu Arg Val Arg Leu Gly Glu Leu Arg	995	1100	1105
Lys Gln His Tyr Val Leu Ala Gly Gly Met Gly Thr Pro Ser Glu	1110	1115	1120
Glu Glu Pro Ser Arg Pro Ser Pro Ala Pro Arg Ser Gly Ala Thr	1125	1130	1135
Lys Lys Pro Pro Leu Ala Gln Lys Pro Ser Ile Ala Pro Arg Thr	1140	1145	1150
Asp Asn Gln Leu Asp Lys Lys Asp Gly Val Tyr Pro Ala Gln Leu	1155	1160	1165
Val Asn Tyr			

(2) INFORMATION FOR SEQ ID NO:12:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 18

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: other DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12:

GAAGATACCC CACCAAAC 18

(2) INFORMATION FOR SEQ ID NO:13:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 35

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: other DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no
(vi) ORIGINAL SOURCE:
(A) ORGANISM: human
(xi)SEQUENCE DESCRIPTION: SEQ ID NO: 13:
GCTTGACAGT GTAGTCATAA AGGTGGCTGC AGTCC 35

(2) INFORMATION FOR SEQ ID NO:14:
(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 24
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear
(ii)MOLECULE TYPE: other DNA
(iii) HYPOTHETICAL: no
(iv) ANTI-SENSE: no
(vi) ORIGINAL SOURCE:
(A) ORGANISM: human
(xi)SEQUENCE DESCRIPTION: SEQ ID NO: 14:
GGACATGTCC AGGGAGTTGA ATAC 24

(2) INFORMATION FOR SEQ ID NO:15:
(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 41
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear
(ii)MOLECULE TYPE: other nucleic acid
(iii) HYPOTHETICAL: no
(iv) ANTI-SENSE: yes
(vi) ORIGINAL SOURCE:
(A) ORGANISM: human
(xi)SEQUENCE DESCRIPTION: SEQ ID NO: 15:
CUACUACUAC UACUAGGCCA CGCGTCGACT AGTACGGGII GGGIIGGGII G 41

(2) INFORMATION FOR SEQ ID NO:16:
(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 516
(B) TYPE: nucleic acid
(C) STRANDEDNESS: double
(D) TOPOLOGY: linear
(ii)MOLECULE TYPE: genomic DNA
(iii) HYPOTHETICAL: no
(iv) ANTI-SENSE: no
(vi) ORIGINAL SOURCE:
(A) ORGANISM: human

(x) FEATURE: exon 1 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:

TCTGTGGAAG	TTTTGGAGGG	GAGAGAGGGG	CAGCTGGATG	CTCTTGGGCC	ACGGTCGCCC	60
CTGATCTCTG	CGCCTCTTCC	TCCTGCTCCG	GGAGAAATAA	TGTTTCCCTG	GGGGATGAAA	120
GCATCTCTTT	GTGCGGGCTT	TAATTGCCAT	GTTGTTGTGC	CAAGGGAGTG	AGTGGCGGCG	180
GGACCAGCAG	CTGGGCACAG	CCAATGCCAG	GCAGTGGTGC	CCACTCCCTC	AGGACGCCCA	240
GCCAGCTGGC	TCCTGGGAGC	GCTGCCCCAC	TCTGCCCCCA	GCTGGGCGCC	TGCAAGGAAC	300
CGACCACCCG	TGGGGCTGGG	GGAGGTTGGC	TGGAGGAGGA	GAAAGGGGCG	GGCTCTGGGA	360
GGGTCTCAGC	CACTCTCAGA	GGCTTATTCA	TCTCATCCTC	CTTTCCTTCC	CCCTTCTTGT	420
TTTTTCAGACT	GTCAGCATCA	ATAAGGCCAT	TAATACGCAG	GAAGTGGCTG	TAAAGGAAAA	480
ACACGCCAGA	AATATCCTTT	GGATGTTGCT	TGGAAG			516

(2) INFORMATION FOR SEQ ID NO:17:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 193

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 2 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17:

TGTTTTCCAT	AACCCCCCT	CACCGTGCAT	ACTGGGCACC	CACCATGAGA	AAGGGGCACA	60
GACCTTCTGG	TCTGTTGTCA	ACCGCTGCC	TCTGTCTAGC	AACCCAGTGC	TCTGCTGGAA	120
GTTCTGCCAT	GTGTTCCACA	AACTCCTCCG	AGATGGACAC	CCGAACGTGA	GTTCCTGGGG	180
CTATGGGGTG	GCA					193

(2) INFORMATION FOR SEQ ID NO:18:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 104

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 3 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:

GTGTTCTTTT	GCCCTGCAG	GTCCTGAAGG	ACTCTCTGAG	ATACAGAAAT	GAATTGAGTG	60
ACATGAGCAG	GATGTGGGTG	AGTTTGAGGA	TGTACTCAGG	AGCC		104

(2) INFORMATION FOR SEQ ID NO:20:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 327
(B) TYPE: nucleic acid
(C) STRANDEDNESS: double
(D) TOPOLOGY: linear
(ii) MOLECULE TYPE: genomic DNA
(iii) HYPOTHETICAL: no
(iv) ANTI-SENSE: no
(vi) ORIGINAL SOURCE:
(A) ORGANISM: human

(x) FEATURE: exon 4 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:

AATTCCTGGC	TGCAGATCTC	TTGACTGTTA	TGTTCTTGTT	GTTGACTCTG	TTTCCCCTCC	60
TCTTCCTAAA	AGGGCCACCT	GAGCGAGGGG	TATGGCCAGC	TGTGCAGCAT	CTACCTGAAA	120
CTGCTAAGAA	CCAAGATGGA	GTACCACACC	AAAGTGAGTC	TCTGCGGACA	GTTCTGCCGC	180
CACCGCCGCC	TCCCCTGCTC	CATCCCTTCA	GCCCCTCCCT	GGGCTCATTT	GTCAGCTCTT	240
TCAGGTAATA	GACAGCCCAG	GCTTCTGAGG	AAGTGTGCAC	ATCATGTACC	CAAGCTGTGA	300
GAGAGGAAAG	CCACCGCCAG	GCCCACG				327

(2) INFORMATION FOR SEQ ID NO:21:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 331
(B) TYPE: nucleic acid
(C) STRANDEDNESS: double
(D) TOPOLOGY: linear
(ii) MOLECULE TYPE: genomic DNA
(iii) HYPOTHETICAL: no
(iv) ANTI-SENSE: no
(vi) ORIGINAL SOURCE:
(A) ORGANISM: human

(x) FEATURE: exon 5 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:

GGGCTCAAGC	AATCCTCCCA	CCTCGGCCCTC	CCAAGTAGCT	GGGACCACAG	GCGTGTGCCA	60
CCACGCCCGG	CTGAGAGAGG	GCTCTTCATG	TCTTCTGCCC	TGACTCCCTT	CCTCTGCCTC	120
CCTTCCAGAA	TCCCAGGTTC	CCAGGCAACC	TGCAGATGAG	TGACCGCCAG	CTGGACGAGG	180
CTGGAGAAAG	TGACGTGAAC	AACTTGTAAG	TGGCTCCTGC	CCTGAGCCCA	GGGAGGGAGA	240
AAGCTTTTGT	GAATGCTGAC	ACTTCTCATA	AGGGTCATGG	AGGGCCTGAT	GGGGGGAGGC	300
CGTGGCTGGG	ATGGGGACCA	AAGCCCCTGG	G			331

(2) INFORMATION FOR SEQ ID NO:22:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 470
(B) TYPE: nucleic acid
(C) STRANDEDNESS: double
(D) TOPOLOGY: linear
(ii) MOLECULE TYPE: genomic DNA
(iii) HYPOTHETICAL: no
(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 6 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 22:

ACTGTCGCTG	TCACGTGTTGA	CTTCACCAGG	CTGCATGGCC	ATAATACCCA	CAAGGCTAAG	60
ACTTGGAGCT	GGAGTTGTGT	GTGTGTTTGC	GCATGCACAT	GAGCATTGGA	GA CTGGAGTA	120
GCGTAGAGCG	TGGGGGAGGG	GACAGGTAAC	AGACCGGCCT	CAGGCTGTGG	AGTGTAAAGCT	180
CTCTTTCCCTC	TTGGGTCCAG	TTTCCAGTTA	ACAGTGGAGA	TGTTTGA CTA	CCTGGAGTGT	240
GAAC TCAACC	TCTTCCAAAC	AGGTGAGTCT	CTTCCCTCCC	GTCTAACCCA	GGCTCTCATG	300
GGAAC TACCT	AATTCCTAGT	CCTCCTCTCC	CTGCAAAGTG	TGCAGCACAA	GGGGTAGGAA	360
AATGGAGACA	TTACACCCCC	ATCTCTGGTC	TCTCCAACCC	TCGTGCAGGG	AGGGACTGAA	420
CCTCTTCAGT	ATTTTCTTTT	TTAAGAGACA	AGGTCTCGGC	CGGGTGCAGT		470

(2) INFORMATION FOR SEQ ID NO:23:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 565

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 7 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23:

TCTTCACCTG	TTTAATGGGG	ATACGTTTAC	CTATCTCATG	GGAGTGTGTG	GAAGGTTAAA	60
TGAATTAGAT	GAGGTAAAGC	ACGCACAGAA	TCGGTCCTTG	GTGTATGTTG	GACCCCTGCC	120
TCTGCCCCCTC	TGAAGAGGCT	GCCTGTAATC	CCCTGGCTCT	ACCACCTTTC	TCCCTCACTT	180
TTATTTCCCTA	GTATTCAACT	CCCTGGACAT	GTCCCGCTCT	GTGTCCGTGA	CGGCAGCAGG	240
GCAGTGCCGC	CTCGCCCCGC	TGATCCAGGT	CATCTTGGAC	TGCAGCCACC	TTTATGACTA	300
CAC TGTCAAG	CTTCTCTTCA	AACTCCACTC	CTGTGAGTAC	CGCGGGCCAG	ATCTTCTTAC	360
ATGAGATTCA	GGCCAGAGGG	AGGATCCCAG	CCTGAGGATG	TCCCCAGAGA	AACGCAGTCC	420
TTCTCAGTGC	CTTTGGCTGT	CTGCTTCTGT	TCCAAAAGGC	CCCGGAGCTT	CTGACCAT TG	480
TGAGGATAAA	AGAGCAGGGC	CCAGGCTTTG	GTGACCC CAG	TAAAGCCCCT	GGCTTGCCAC	540
TCTTGCGTCC	AGTGTTACAG	GATCT				565

(2) INFORMATION FOR SEQ ID NO:24:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 233

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 8 of HIP1

(xi)SEQUENCE DESCRIPTION: SEQ ID NO: 24:

GGGACAGCTC	TAGGCCAGTC	GTGGCCCCCTG	GCAGTGCTGG	CCACATGCCC	CAGGGTAGCT	60
GGGCCCCCTCC	CCCTCGAGAG	CCCCGCTGTG	GCTTCCCTGC	CCTCTGGTCC	CCCTCCCCCTC	120
TCACACTCTT	TCCAATTTCT	TCCAGGCCCTC	CCAGCTGACA	CCCTGCAAGG	CCACCGGGAC	180
CGCTTCATGG	AGCAGTTTAC	AAAGTAAGTG	GTTCAAGTAA	CAGGAATGGA	GGT	233

(2) INFORMATION FOR SEQ ID NO:25:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 578

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii)MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exons 9 and 10 of HIP1

(xi)SEQUENCE DESCRIPTION: SEQ ID NO: 25:

TGAATCCCAG	CACCATGGAG	TTTATCTCCT	TGACAGCCTG	TGCCTTTGGG	CTGGGGAGGG	60
GGCAGGAAAG	CCAGGTGGCT	GCTCTGTCCC	CTACATGGGG	CTGATGAAGA	CACCCAGCAC	120
CCCTCAGGTC	CTTCTCCACC	CCTAGGTGTA	AAGATCTGTT	CTACCGCTCC	AGCAACCTGC	180
AGTACTTCAA	GCGGCTCATT	CAGATCCCCC	AGCTGCCTGA	GGTAAGCATG	CCCAACCACA	240
CACCTCGGC	ACTGCAGAGG	CCCCAGGTAC	TCTCTTAAGG	GCCGGCGGGG	CCTGGCAAGC	300
AAGCACTATT	TGAGGATGTG	TCTCCGTCTT	CAGAACCCAC	CCAACTTCCT	GCGAGCCTCA	360
GCCCTGTCTG	AACATATCAG	CCCTGTGGTG	GTGATCCCTG	CAGAGGCCTC	ATCCCCCGAC	420
AGCGAGCCAG	TCCTAGAGAA	GGATGACCTC	ATGGACATGG	ATGCCTCTCA	GCAGGTGAGG	480
ACCACTTGGG	AGAGAAACTT	GGCCTTTCCT	CTCACCTGCA	AGTACAGGGG	AGAGGCTGGG	540
GGAGACCCTG	GCCAAAGCCC	ATTGACTCTA	ACCAGGTT			578

(2) INFORMATION FOR SEQ ID NO:26:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 390

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii)MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 11 of HIP1

(xi)SEQUENCE DESCRIPTION: SEQ ID NO: 26:

AAAAAAATTT	AAAAAATTAA	ACAGGTCTGA	ACCGTTTAAT	TCGAGAAAGG	GGGCATTCTC	60
CCATATCACT	CAACTGACCC	ACACACAGAA	TTCTCTGGCT	CTCTGACTTA	TTCTCACTCC	120
TTTTTGGTCA	ACCACAGAAT	TTATTTGACA	ACAAGTTTGA	TGACATCTTT	GGCAGTTCAT	180
TCAGCAGTGA	TCCCTTCAAT	TTCAACAGTC	AAAATGGTGT	GAACAAGGAT	GAGAAGTGAG	240
TCCAAGCTGG	GTTCAAGCAG	ATGGTTCAGG	AGCTAAGTTA	AGCCATGGTC	TGCCCTCAAAA	300
CACTAACCAA	AGAGGAATTC	TTAATGATAC	TGGGGCTTCT	TAGATACAGA	ACATCTTGAA	360
GGGTTGGGGG	CAATGGCTTA	TGCCTGTAAT				390

(2) INFORMATION FOR SEQ ID NO:27:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 547

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 12 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:

AAAATCAATA	ACCATGGATT	TATGAGTATT	AGATTAGTAT	CTGGTAACAT	TTAGAGTATA	60
ATTTATGGCA	TTTCAAAGAA	TTGTCCCCAA	ATTAATACCA	GCTTTTAATT	TCCTCCCCTG	120
AGCTCACAAT	TAAAAACAGA	GGGATAGAAG	CACATATGAAA	GCAAACATCAT	TCCCCTTCTC	180
TTCCAGGGA	CCACTTAATT	GAGCGACTAT	ACAGAGAGAT	CAGTGGATTG	AAGGCACAGC	240
TAGAAAACAT	GAAGACTGAG	GTATAACTTG	GATCTGCTCT	GCCTTTGCGC	TTACCAAAAA	300
CACGGTAGAT	TTGAATGTTA	AATTTGCATC	ACACTAGCCA	GGCACAGTGG	CTCACACCTG	360
TAATCCTAGC	ACTTTGGGAG	GCCAAGGCAG	GAGGATTACC	TGAGGTCGGG	AGTTCGAGAC	420
CAGCCTGGGC	AACAGGTGA	AACCCCGTC	TTCAATAAAA	ATGCAATAAT	TAGCCGGGTG	480
TGTTGGCAGG	CACCTGTAAT	CCCAGCTACT	CGGGAAGCTG	AGGCATGAGA	ATTGCTTGAA	540
CTTGGA						547

(2) INFORMATION FOR SEQ ID NO:28:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 436

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 13 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 28:

CCCCCAGCCA	CTCTAAAGAG	GACCACAATT	CCCCGGCCAT	CATCCCCTGT	TATTGTTGTT	60
GATTGAGGGG	CTCCTAATGA	CCAGATGGTC	CAACCTCCT	GGGACGTGGA	GAGTTGACTT	120
AGGGGAATCA	GGTATTTACT	TGGAAGCATG	GTAGGACCCG	CTTCTCCGGC	CCATGCCCGT	180
GACCCGTGGC	AGTGGGCGGT	TGGCCTCATG	ACCGGAGTCC	CCCCACAGAG	CCAGCGGGTT	240
GTGCTGCAGC	TGAAGGGCCA	CGTCAGCGAG	CTGGAAGCAG	ATCTGGCCGA	GCAGCAGCAC	300
CTGCGGCAGC	AGGCGGCCGA	CGACTGTGAA	TTCTGCGGG	CAGAACTGGA	CGAGCTCAGG	360
AGGCAGCGGG	AGGACACCGA	GAAGGCTCAG	CGGAGCCTGT	CTGAGATAGA	AAGTGAGCGG	420
TGGGTGGGGG	CGGGG					436

(2) INFORMATION FOR SEQ ID NO:29:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 469
(B) TYPE: nucleic acid
(C) STRANDEDNESS: double
(D) TOPOLOGY: linear
(ii) MOLECULE TYPE: genomic DNA
(iii) HYPOTHETICAL: no
(iv) ANTI-SENSE: no
(vi) ORIGINAL SOURCE:
(A) ORGANISM: human
(x) FEATURE: exon 14 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:

GACTTGAGCC	CAAGGAGGTC	AAGGCTGCAG	TGAACAGTGA	TTGTGCCACT	GCACCCCAGC	60
CTGGGTGACA	GAGCAAGACT	GTCTCAAAAC	AAAACAAGGA	GGACCTTCTA	GGGACCCTGG	120
CTCATTGCAA	GGAAGGCAAG	GGTCCCTGCT	AGGTTAGACT	CCTCACCTTG	GTCTTTTACA	180
ATACAGGGAA	AGCTCAAGCC	AATGAACAGC	GATATAGCAA	GCTAAAGGAG	AAGTACAGCG	240
AGCTGGTTCA	GAACCACGCT	GACCTGCTGC	GGAAGGTAAG	ACCCTCAGCC	CCTGTCACCA	300
TCCTGCAGGC	CCTGCACCTC	TAGGGAGAGA	GCGGCTCAGG	CCTGTGGCTT	CCCCGGGGCC	360
AGCAACCCCT	ACATTGATCT	CTAAGGCATT	GCCGTCATCT	CGGGAACCAC	ACCTTTTCAG	420
GCTTCCTTGC	CTCTGTGTCT	TGGGCTGTGT	CCTGGGTGCC	AATCCCATG		469

(2) INFORMATION FOR SEQ ID NO:30:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 359
(B) TYPE: nucleic acid
(C) STRANDEDNESS: double
(D) TOPOLOGY: linear
(ii) MOLECULE TYPE: genomic DNA
(iii) HYPOTHETICAL: no
(iv) ANTI-SENSE: no
(vi) ORIGINAL SOURCE:
(A) ORGANISM: human
(x) FEATURE: exon 15 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:

GGGTAGGAAA	GTGATTCCTG	TGTCTGACTC	TAGGGCACGC	ACAGCCTGAG	TATGATTGTC	60
CTAGAAGGAG	GATGTCCTCT	AAGCCTGGGA	TCTCCTGGTT	CAAGACACTG	TTCTTCTTTT	120
GCAGAATGCA	GAGGTGACCA	AACAGGTGTC	CATGGCCAGA	CAAGCCCAGG	TAGATTTGGA	180
ACGAGAGAAA	AAAGAGCTGG	AGGATTCGTT	GGAGCGCATC	AGTGACCAGG	GCCAGCGGAA	240
GGTGAGTGGG	ACGAGGAGCA	CTCGGGAAAT	GAGGGAGGGG	GCTGTTGAGT	TGGTGGCGGG	300
GGCTTTGTGG	CCTTCTGCTC	CATGGGCAGT	TCTGTGGGTC	GTTTGGCATC	ACACAGCAG	359

(2) INFORMATION FOR SEQ ID NO:31:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 209
(B) TYPE: nucleic acid
(C) STRANDEDNESS: double
(D) TOPOLOGY: linear
(ii) MOLECULE TYPE: genomic DNA
(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 16 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:

GTTGATCGCT	TGGGACGTTT	TTACATTTTT	ATATTCTTTG	TCAGTGTAC	CCAGATCAGA	60
GTCCCTCTGT	TTTTCTTCTC	TTTCAGACTC	AAGAACAGCT	GGAAGTTCTA	GAGAGCTTGA	120
AGCAGGAACT	TGCCACAAGC	CAACGGGAGC	TTCAGGTTCT	GCAAGGCAGC	CTGGAAACTT	180
CTGCCCAGGT	AAATACCTCC	TTTTTTTTTT				209

(2) INFORMATION FOR SEQ ID NO:32:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 485

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 17 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:

CCCCCACTGC	AATCAGTGTG	TCCCCGGGAG	GGAATCAGAG	TGGCAGGTTA	AAGAGCCATC	60
ACCTTCCAG	TCCTTGCAAC	CCGGTGGTGG	GTTGGACCTC	TGGGAAGTAG	GGACTGTTTA	120
ACTCAACCAG	CGTCTCCCTC	TTTCCTTGTC	GTCACCTTTG	CAGTCAGAAG	CAAACCTGGGC	180
AGCCGAGTTC	GCCGAGCTAG	AGAAGGAGCG	GGACAGCCTG	GTGAGTGGCG	CAGCTCATAG	240
GGAGGAGGAA	TTATCTGCTC	TTCGGAAAGA	ACTGCAGGAC	ACTCAGCTCA	AACTGGCCAG	300
CACAGAGGGT	CACGGACATG	GACACGAGCG	AGCACCTGTG	AATTCCCACC	GAGGGCCTCT	360
GCGCATGCAC	GGAGGCTGGG	AGGACCCCGG	GGCTGCTGAG	AAGGGGTTTG	GGGCCTTGGC	420
CTGATTGTGC	AGACATTCTG	TAGGTGTAAT	GCCAGCAGGC	CCTGCATTGC	CTGCAGAGTC	480
CATGA						485

(2) INFORMATION FOR SEQ ID NO:33:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 468

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 18 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:

TTACTGGCTT	GGACCTCATT	GGCCATGACT	TGAGCTAAGA	TGCTAAGAGC	CCCAGCCAGG	60
TCATCCTGCT	CAGGTTTATT	ATGGAGTCTA	GGGCAGACTC	TCACCTCCCT	GGACCATTTT	120

TAGAATCTAT	GTGCCAGCTT	GCCAAAGACC	AACGAAAAAT	GCTTCTGGTG	GGGTCCAGGA	180
AGGCTGCGGA	GCAGGTGATA	CAAGACGCCC	TGAACCAGCT	TGAAGAACCT	CCTCTCATCA	240
GCTGCGCTGG	GTCTGCAGGT	ACACTTGCAA	TTGCCCAGCT	GGCAGGGGCC	AGGTCCTTAC	300
AGCCTGAGAC	TCTGTTGATG	TTGAATCTCA	TGTGAGACTT	AGCTCAGGGG	CTCTCAGCCC	360
AGCAGCATGT	CAGCATTACC	TTAGGGGCGC	CCAGGCCCCA	TCCTAGATCA	GTTACATGTG	420
GAAACTCTGT	GCATTAGTGC	CTATACACTA	GTATTTTAGT	ATTTTCTT		468

(2) INFORMATION FOR SEQ ID NO:34:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 393

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 19 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34:

CACTAGTAAG	CTCCTCCATT	CAGTGCTTAA	TTAACGAGGA	TGAAGCCAGC	TATGAGAACT	60
TGCTCTGACC	TTGCCCTGTG	TTCCCTCTCA	CAGATCACCT	CCTCTCCACG	GTACATCCA	120
TTTCCAGCTG	CATCGAGCAA	CTGGAGAAAA	GCTGGAGCCA	GTATCTGGCC	TGCCCAGAAG	180
GTAAGAATGG	CCAAGGACAG	TCTCTGTCTG	CTAGTGATGG	CCAGACAGGG	TTCAGAAGCA	240
CCTGAATGCG	GGGATAGTGA	CAGGTCCCTC	TGCATCAAGA	AAGGCATGTA	GGCAACTCAT	300
ACAAGAAAGG	CATGTAGGCA	ACTCATAAAA	CGGGAGGAGA	GGGTATGAAA	GTGTCACCAT	360
CAACCAGACC	TGAGAAACTT	CTCTTTCCAA	TCC			393

(2) INFORMATION FOR SEQ ID NO:35:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 421

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 20 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:

GGCCTGCCCA	GAAGGTAAGA	ATGGCCAAGG	ACAGTCTCTG	TCGGCTAGTG	ATGGCCAGAC	60
AGGGTTTACA	AGCACCTGAA	TGCGGGGATA	GTGACAGGTC	CCTCTGCATC	AAGAAAGGCA	120
TGTAGGCAAC	TCATACAAGA	AAGGCATGTA	GGCAACTCAT	AAAACGGGAG	GAGAGGGTAT	180
GAAAGTGTCA	CCATCAACCA	GACCTGAGAA	ACTTCTCTTT	CCAATCCTGG	CAGACATCAG	240
TGGACTTCTC	CATTCATATA	CCCTGCTGGC	CCACTTGACC	AGCGACGCCA	TTGCTCATGG	300
TGCCACCACC	TGCCCTCAGAG	CCCCACCTGA	GCCTGCCGAC	TGTGAGTACT	GGGGCATGAG	360
GGGCTGTTCA	TGGACCAGGG	GAGCAGGGGG	CCTTTAAAAG	TCTCTGTTGG	GCCGGGCGCA	420
G						421

(2) INFORMATION FOR SEQ ID NO:36:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 498

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 21 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36:

AGGCCGAGGC	AGGAGAATCG	CTTGAACCTCA	GGAGGCGGAG	TTTGCAGTGA	GCCGAGATGG	60
CGCCACTGCA	CTCCAGCCTG	GGCAACAAGA	GCGAGACTCC	ATCTCAAAAA	AAAAGTGTCT	120
ATTGCCTTGT	ATCTCCAGCA	CTGACCGAGG	CCTGTAAGCA	GTATGGCAGG	GAAACCCCTCG	180
CCTACCTGGC	CTCCCTGGAG	GAAGAGGGAA	GCCTTGAGAA	TGCCGACAGC	ACAGCCATGA	240
GGAATGCCT	GAGCAAGATC	AAGGCCATCG	GCGAGGTACT	TGGAGTAGTA	TCATTGAGGA	300
GCATTGTTAT	TCTTCTGGGT	GTGCGTGCTG	GTGAATGGCC	AGGGAATCGG	TGATGTTCTG	360
AGCTAGTTCT	TTCTGCACTT	AGAACTTGAT	TCTAGAAAGA	GATTGTTAAA	ATTGGAAAAT	420
CTGGCCGGGT	GCAGTGATTT	ATGCGTGTA	TCCGAGCACT	TTGGGAGGCC	GAGTCAGGAG	480
GATCACTTGA	GGCTAGAC					498

(2) INFORMATION FOR SEQ ID NO:37:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 427

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 22 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 37:

CCCTGTGGCT	TGCAGAAGGT	GTTTGCTGGG	TGGCCTCCTG	CCTTGCCATC	TTGTAAGGGT	60
TACAGATGGC	AGAGGAGAAG	AGACAGGAGG	CCCCAAGGTC	AGTTCAGCCT	TTGTGATGTG	120
TTACAGGAG	CTCCTGCCCA	GGGGACTGGA	CATCAAGCAG	GAGGAGCTGG	GGGACCTGGT	180
GGACAAGGAG	ATGGCGGCCA	CTTCAGCTGC	TATTGAAACT	GCCACGGCCA	GAATAGAGGT	240
AGGAGGTTCC	TGCAGGATCT	CCTGAAACGA	TGCCTTTGCA	GCTGCCCTTC	TGCAACACTG	300
CTCATTAAC	ATGTCACAGT	CGTTCATTAA	GGCCATGGCA	ACCCCTAAG	ACAGAAACCA	360
GAATTTGCCA	GGCACAGTGG	CTCATGCCTG	TAACCCAGC	ACCTTGGGAG	GATCACTTGA	420
GTCCAGG						427

(2) INFORMATION FOR SEQ ID NO:38:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 367

(B) TYPE: nucleic acid
(C) STRANDEDNESS: double
(D) TOPOLOGY: linear
(ii) MOLECULE TYPE: genomic DNA
(iii) HYPOTHETICAL: no
(iv) ANTI-SENSE: no
(vi) ORIGINAL SOURCE:
(A) ORGANISM: human
(x) FEATURE: exon 23 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 38:

CCCCCTGAAT	AGGTTAGAGT	CTGGATTCTT	TTCTGACTCT	CTCAAGAATG	TGGGCAGGGA	60
CTTGGGGACT	TCCAGATTCA	GGTTTCCCAG	CTACCACACG	ATGTTGGACT	GAAAGTATAG	120
TAAGACATTA	GTGGATCCTT	AATATTCAAG	GCACATTTAG	AAACCATGCT	TCTTTTTTAC	180
AGGAGATGCT	CAGCAAATCC	CGAGCAGGAG	ACACAGGAGT	CAAATTGGAG	GTGAATGAAA	240
GGTCGGTCTG	AGCGGCATGG	TGGGACCTAG	GGGAGCAGGA	TCTGTCTTCC	TGACATTGGT	300
CTATACTTTG	CATACTTATT	AGGGAATTAG	AGGAGAGCAG	TAGCAGCCAC	GGGGAAGGGC	360
TGAGTTG						367

(2) INFORMATION FOR SEQ ID NO:39:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 502

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 24 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:

CCCCGCAGAA	TGTTCCAGCA	ACCTCAGCAC	CCTTCTTACC	TCCCTTTCCC	ATTCCAAGCT	60
TGCCTTTGGC	TAGGAGTGGG	GAAGAGAACC	GTCGTGTTCA	TTGATCTTGG	ATCTTGATCT	120
CAGTGTATCC	TCGACTTGTT	TGTTTGGCAG	GATCCTTGGT	TGCTGTACCA	GCCTCATGCA	180
AGCTATTGAG	GTGCTCATCG	TGGCCTCTAA	GGACCTCCAG	AGAGAGATTG	TGGAGAGCGG	240
CAGGGTGAGC	GTGGGTGTGG	GCCCTGGGCA	GGAAGAGGAG	GCATCGGTGA	CAGACTCCCG	300
CTCCAACGGA	CTCTGTGATG	CTGCCGTCTT	ACTCTGTGTG	TCCACCTGAG	TACAGAGCAG	360
CCACTCCTGT	AGATATCAGC	AGAGGCCCTG	GGGAGAAGTC	AGAGCTCCAG	GACCTCCCCA	420
GAGGGTGGCC	AGGCATGTGT	CCCAACTCCA	GCTCCCTTCG	CACAGGCAGA	CATTGTTGGA	480
ACTTGCTGTG	GGAGCCCTTT	TT				502

(2) INFORMATION FOR SEQ ID NO:40:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 437

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 25 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 40:

TTTTGGTCTC	TGAATCTTCT	TCTTTTTTGT	AAAATGGGAA	TACTAATGCT	TATGTCTCAG	60
AGTTACTATG	AGGATGATTT	GGGATAATAT	ATGTATAAAA	GCACCTGCCA	TATAGTACAT	120
GCTCAATAAA	AGGTGGCTAT	TACTATTTTT	TATTTCCCTA	GGGTACAGCA	TCCCCTAAAG	180
AGTTTTATGC	CAAGAACTCT	CGATGGACAG	AAGGACTTAT	CTCAGCCTCC	AAGGCTGTGG	240
GCTGGGGAGC	CACTGTCATG	GTGTAAGTAT	CTATTGGTAC	CAAGGGTCCT	CCCATGACCC	300
CTCTTCCATT	GATCCACTCC	AAACAATAGC	TAAGGAGGGA	AAAAAAAATC	TGTCCCTTAG	360
AAATAAACTA	TTGATCAGGA	AGTCAATAGG	ACCGAGTTTA	CAAGGGAGCC	TGGCTCTCCC	420
AGGGGACACA	GGGCAGG					437

(2) INFORMATION FOR SEQ ID NO:41:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 351

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 26 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:

GGGAGCCTGG	CTCTCCAGG	GGACACAGGG	CAGGCAGCCT	CCCCTCCCTG	TTAGCCAAG	60
GGCGATGGGG	TGGTCTGGAG	GTGGGATTGT	GGAGGAGTTG	CAGCTCATTT	GCCCGTAACC	120
TAGTCCCTCT	TGTCGTTTTC	CATCAGGGAT	GCAGCTGATC	TGGTGGTACA	AGGCAGAGGG	180
AAATTTGAGG	AGCTAATGGT	GTGTTCTCAT	GAAATTGCTG	CTAGCACAGC	CCAGCTTGTG	240
GCTGCATCCA	AGGTAGGACC	TGGCTGGACC	TCCTAGGACG	CTGGAAGGCC	TGGTTAGAGA	300
GTACTAGGCT	AGGTTAAAGA	GTACTTGGCT	GCGTTAGGCA	GTACTTGGCT	G	351

(2) INFORMATION FOR SEQ ID NO:42:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 418

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 27 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:

CTTTTATAT	GATAGATATG	TCAGGAGCTG	ACTATAGTCA	GCAGATTTTG	AGAAGCTGAT	60
TGGTGATTGC	CGTTTGGCCC	ACATATGTTT	GCTAAGAACC	ATCAGAGCAA	TTATCTGATT	120
CAGTCCTTGT	TGCTCTAGGT	GTTGTATGAA	CCTAAATCTG	CTTTGTCTCTG	GTAGGTGAAA	180

GCTGATAAGG	ACAGCCCCAA	CCTAGCCCAG	CTGCAGCAGG	CCTCTCGGGG	AGTGAACCAG	240
GCCACTGCCG	GCGTTGTGGC	CTCAACCATT	TCCGGCAAAT	CACAGATCGA	AGAGACAGGT	300
AGCCTTTCCA	AAGGGACCCT	TTTCTTACCC	ACCCTGTTGA	GCTCTTCTCT	GCATCCTTCC	360
CTGTGATCCC	AACCAAATCC	CACAGGACTG	TGTCTAAATT	CTTTCATATT	TTTCATCT	418

(2) INFORMATION FOR SEQ ID NO:43:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 279

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 28 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:

TTTCCACAGA	GCATTGGCAT	TGGCTGCCTC	TCAGGTGCCA	GTCAGCCAGG	GTAGAATTTG	60
ATGAGACCTT	CTTGTTTCCA	TCCTTGCAGA	CAACATGGAC	TTCTCAAGCA	TGACGCTGAC	120
ACAGATCAAA	CGCCAAGAGA	TGGATTCTCA	GGTTAGGGTG	CTAGAGCTAG	AAAATGAATT	180
GCAGAAGGAG	CGTCAAAAAC	TGGGAGAGCT	TCGGAAAAAG	CACTACGAGC	TTGCTGGTGT	240
TGCTGAGGGC	TGGGAAGAAG	GTAAGCTGAC	TCAAAGGAT			279

(2) INFORMATION FOR SEQ ID NO:44:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 3715

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 29 and partial cds of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 44:

AACATAAATT	ATCATTGTCT	TTTAGGAACA	GAGGCATCTC	CACCTACACT	GCAAGAAGTG	60
GTAACCGAAA	AAGAATAGAG	CCAAACCAAC	ACCCCATATG	TCAGTGTAAG	TCCTTGTTAC	120
CTATCTCGTG	TGTGTTATTT	CCCCAGCCAC	AGGCCAAATC	CTTGGAGTCC	CAGGGGCAGC	180
CACACCACTG	CCATTACCCA	GTGCCGAGGA	CATGCATGAC	ACTTCCCAAA	GACTCCCCTC	240
ATAGCGACAC	CCTTCTGTGT	TGGACCCATG	GTCATCTCTG	TTCTTTTCCC	GCCTCCCTAG	300
TTAGCATCCA	GGCTGGCCAG	TGCTGCCCCAT	GAGCAAGCCT	AGGTACGAAG	AGGGGTGGTG	360
GGGGGCAGGG	CCACTCAACA	GAGAGGACCA	ACATCCAGTC	CTGCTGACTA	TTTGACCCCC	420
ACAACAATGG	GATATCCTTAA	TAGAGGAGCT	GCTTGTTGTT	TGTTGACAGC	TTGGAAAGGG	480
AAGATCTTAT	GCCTTTTCTT	TTCTGTTTTC	TTCTCAGTCT	TTTCAGTTTC	ATCATTTGCA	540
CAAACCTTGT	AGCATCAGAG	GGCTGATGGA	TTCCAAACCA	GGACACTACC	CTGAGATCTG	600
CACAGTCAGA	AGGACGGCAG	GAGTGTCTCT	GCTGTGAATG	CCAAAGCCAT	TCTCCCCCTC	660
TTTGGGCAGT	GCCATGGATT	TCCACTGCTT	CTTATGGTGG	TTGGTTGGGT	TTTTTGGTTT	720
TGTTTTTTTT	TTTTAAGTTT	CACTCACATA	GCCAACTCTC	CCAAAGGGCA	CACCCCTGGG	780
GCTGAGTCTC	CAGGGCCCCC	CAACTGTGGT	AGCTCCAGCG	ATGGTGCTGC	CCAGGCCTCT	840

CGGTGCTCCA	TCTCCGCTC	CACACTGACC	AAGTGCTGGC	CCACCCAGTC	CATGCTCCAG	900
GGTCAGGCGG	AGCTGCTGAG	TGACAGCTTT	CCTCAAAAAG	CAGAAGGAGA	GTGAGTGCCT	960
TTCCCTCCTA	AAGCTGAATC	CCGGCGGAAA	GCCTCTGTCC	GCCTTTACAA	GGGAGAAGAC	1020
AACAGAAAAG	GGGACAAGAG	GGTTCACACA	GCCCAGTTCC	CGTGACGAGG	CTCAAAAAC	1080
TGATCACATG	CTTGAATGGA	GCTGGTGAGA	TCAACAACAC	TACTTCCCTG	CCGGAATGAA	1140
CTGTCCGTGA	ATGGTCTCTG	TCAAGCGGGC	CGTCTCCCTT	GGCCAGAGA	CGGAGTGTGG	1200
GAGTGATTCC	CAACTCCTTT	CTGCAGACGT	CTGCCCTGGC	ATCCTCTTGA	ATAGGAAGAT	1260
CGTTCCACTT	TCTACGCAAT	TGACAAACCC	GGAAGATCAG	ATGCAATTGC	TCCCATCAGG	1320
GAAGAACCCT	ATACTTGGTT	TGCTACCCTT	AGTATTTATT	ACTAACCTCC	CTTAAGCAGC	1380
AACAGCCTAC	AAAGAGATGC	TTGGAGCAAT	CAGAACTTCA	GGTGTGACTC	TAGCAAAGCT	1440
CATCTTTCTG	CCCGGCTACA	TCAGCCTTCA	AGAATCAGAA	GAAAGCCAAG	GTGCTGGACT	1500
GTTACTGACT	TGGATCCCAA	AGCAAGGAGA	TCATTTGGAG	CTCTTGGGTC	AGAGAAAATG	1560
AGAAAGGACA	GAGCCAGCGG	CTCCAACCTC	TTTCAGCCAC	ATGCCCCAGG	CTCTCGCTGC	1620
CCTGTGGACA	GGATGAGGAC	AGAGGGCACA	TGAACAGCTT	GCCAGGGATG	GGCAGCCCAA	1680
CAGCACTTTT	CCTCTTCTAG	ATGGACCCCA	GCATTTAAGT	GACCTTCTGA	TCTTGGGAAA	1740
ACAGCGTCTT	CCTTCTTTAT	CTATAGCAAC	TCATTTGGTG	TAGCCATCAA	GCACTTCCCA	1800
GGATCTGCTC	CAACAGAATA	TTGCTAGGTT	TTGCTACATG	ACGGGTGTGT	AGACTTCTGT	1860
TTGATCACTG	TGAACCAACC	CCCATCTCCC	TAGCCACCCC	CCCTCCCCAA	CTCCCTCTCT	1920
GTGCATTTTC	TAAGTGGGAC	ATTCAAAAAA	CTCTCTCCCA	GGACCTCGGA	TGACCATACT	1980
CAGACGTGTG	ACCTCCATAC	TGGGTTAAGG	AAGTATCAGC	ACTAGAAATT	GGGCAGTCTT	2040
AATGTTGAAT	GCTGCTTTCT	GCTTAGTATT	TTTTTGTATT	AAGGCTCAGA	AGGAATGGTG	2100
CGTGGCTTCC	TTGTCCCAGT	TGTGGCAACT	AAACCAATCG	GTGTGTTCTT	GATGCGGGTC	2160
AACATTTCCA	AAAGTGGCTA	GTCTCACTT	CTAGATCTCA	GCCATTCTAA	CTCATATGTT	2220
CCCAATTACC	AAGGGGTGGC	CGGGCACAGT	GGCTCACGCC	TGTAATCCCA	GCACTTTGAG	2280
AGGCTGAGGT	GGTAGGATCA	CCTGAGGTCA	GGAGTTCAAG	ACCAGCCTGT	CCAACATGGT	2340
GAAACCCCCA	TCTCTACTAA	AAATACCAA	AATTAGCCGA	GCGTAGTGAC	GGGTGCCCGT	2400
AATCCCAGCT	ACTCAGGAGG	CTGAGACAGG	AGAATCACCT	GAACCCAGAG	GGCAGAGGTT	2460
GCAGTGAGCT	GAGATCACGC	CATTGTACTC	CAGCCTGGGC	AACAAGAGCA	AAACTCCGTC	2520
TCAAAAAAAA	AAAAAATTA	CAAATGGGGC	AAACAGTCTA	GTGTAATGGA	TCAAATTAAG	2580
ATTCTCTGCC	CAGCCGGGCA	CAGTGGCGCA	TGCCTGTAAT	CCCAGAACTT	TGGGAGGCCA	2640
AGACGGGATG	ATTGCTTGAG	CTCAGGAGTT	TGAGACCAGG	CTGGGCATCA	TAGCAAGACC	2700
TCATCTCTAC	TAAATTTCAA	AAACAAAATT	AGCCGGGCAT	GATGGTGCAT	GCCTGTAGTC	2760
TCAGCTAGTT	GGGGAGCTAA	GGTGGGAGAA	TTGCTTGAGC	TTGGGAAGTC	GAGGCTGCAG	2820
TCAGCCCTGA	TTGTGCCAGT	GCACTCCGGC	CTGGGTGACA	GAGTGAGACC	CGTGCTCAAA	2880
AAAAAAAAGA	TTCTGTGTCA	GAGCCCAGCC	CAGGAGTTTG	AGGCTGCAAT	GAGCCATGAT	2940
TTCCCACTGC	ACTCCAGCCT	GAGTGACAGA	GCGAGACTCC	ATCTCTTTAA	AAACAAACAA	3000
AAAATTATCT	GAATGATCCT	GTCTCTAAAA	AGAAGCCACA	GAAATGTTTA	AAAACCTTCAT	3060
CGACTTAGCC	TGAGTCATAA	CGGTAAAGAA	AGCACTTAAA	CAGAAGCAGA	GGCTAATTCA	3120
GTGTCACATG	AGGAAGTAGC	TGTCAGATGT	CACATAATTA	CTTTCGTAAT	AGCTCAGATT	3180
AGAATGGCTA	CCCCATTCTC	TAGACAAAAT	CAAATTGTCC	TATTGTGACT	CTTCTAAAAA	3240
TGAAGATGAA	GAGCTATTTA	ATGACACACC	TTGGATTAAA	ACGGGAATCA	CATCTTAAAG	3300
CTAAAAATGA	ACCTGCAAGC	CTTCTAAATG	AGTCACTGAG	CATCACTAGT	GACAAGTCTC	3360
GGGTGAGCGT	AAATGGGTCA	TGACAAGATG	GGACAGCAAC	AAAATCATGG	CTTAGGATCG	3420
ACAAGAAGTT	AAAAAACAGC	TGCATCTGTT	ACTTAAGTTT	GTAAGACAGT	GCCCTGAGAC	3480
CTCTAGAGAA	AAGATGTTTG	TTTACATAAG	AGAAAGAAGG	CCAGACATGG	TGTCTCACAC	3540
GTTTAAATCCC	AGCACTTTGG	GAGGCAGGGG	CGGGTGGATC	ACCTGAGGTC	AGGAGTTCAA	3600
GACTAGCCTG	GCCAACATGG	TGAAACCCCG	TCTCTACTAA	AAATACAAAA	ATTAGCCGGG	3660
CATGGTGGCA	GGCGCCTATA	ATCCCAGCTA	CTGGGGAGGC	TGAGGCAGGA	GAATC	3715



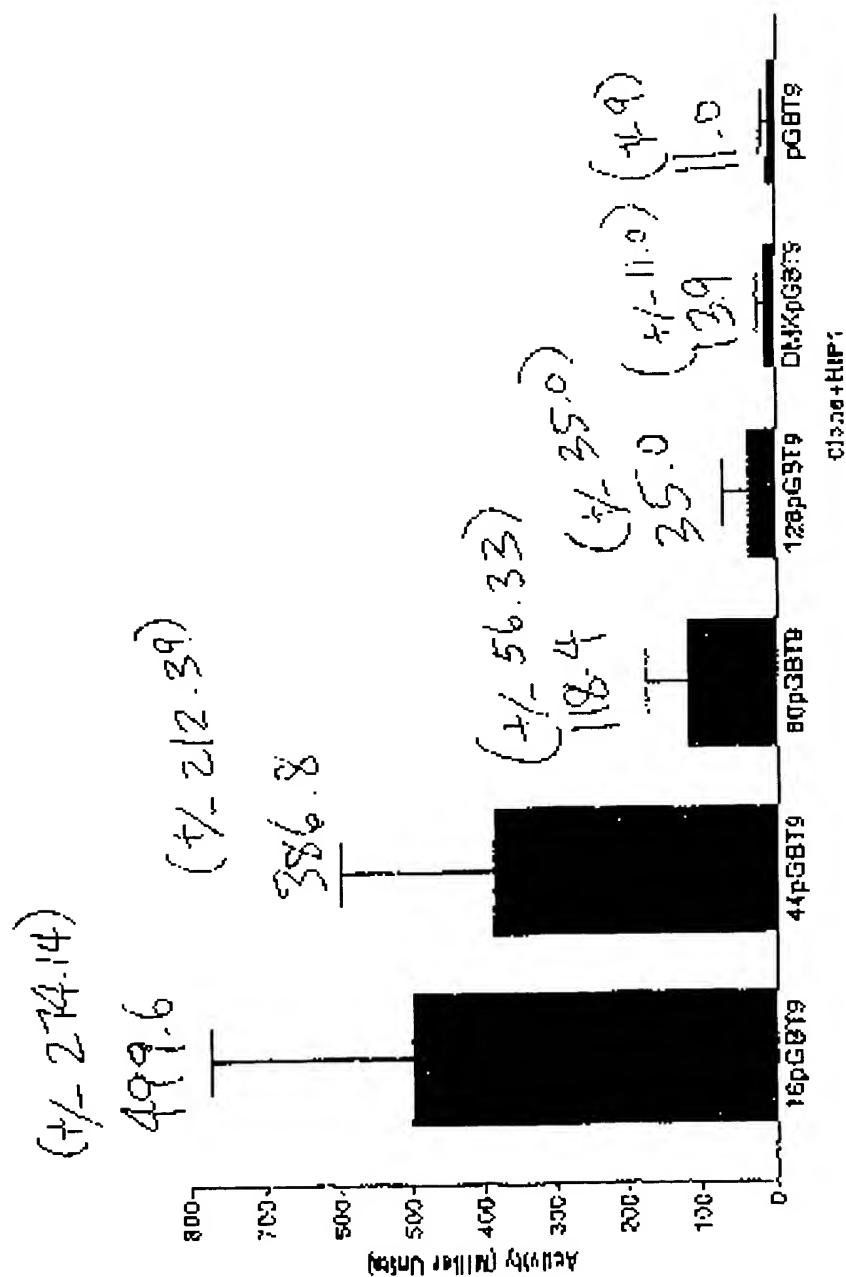


Fig. 1

Fig 2

HIP1 Clones: Nucleotide Alignment

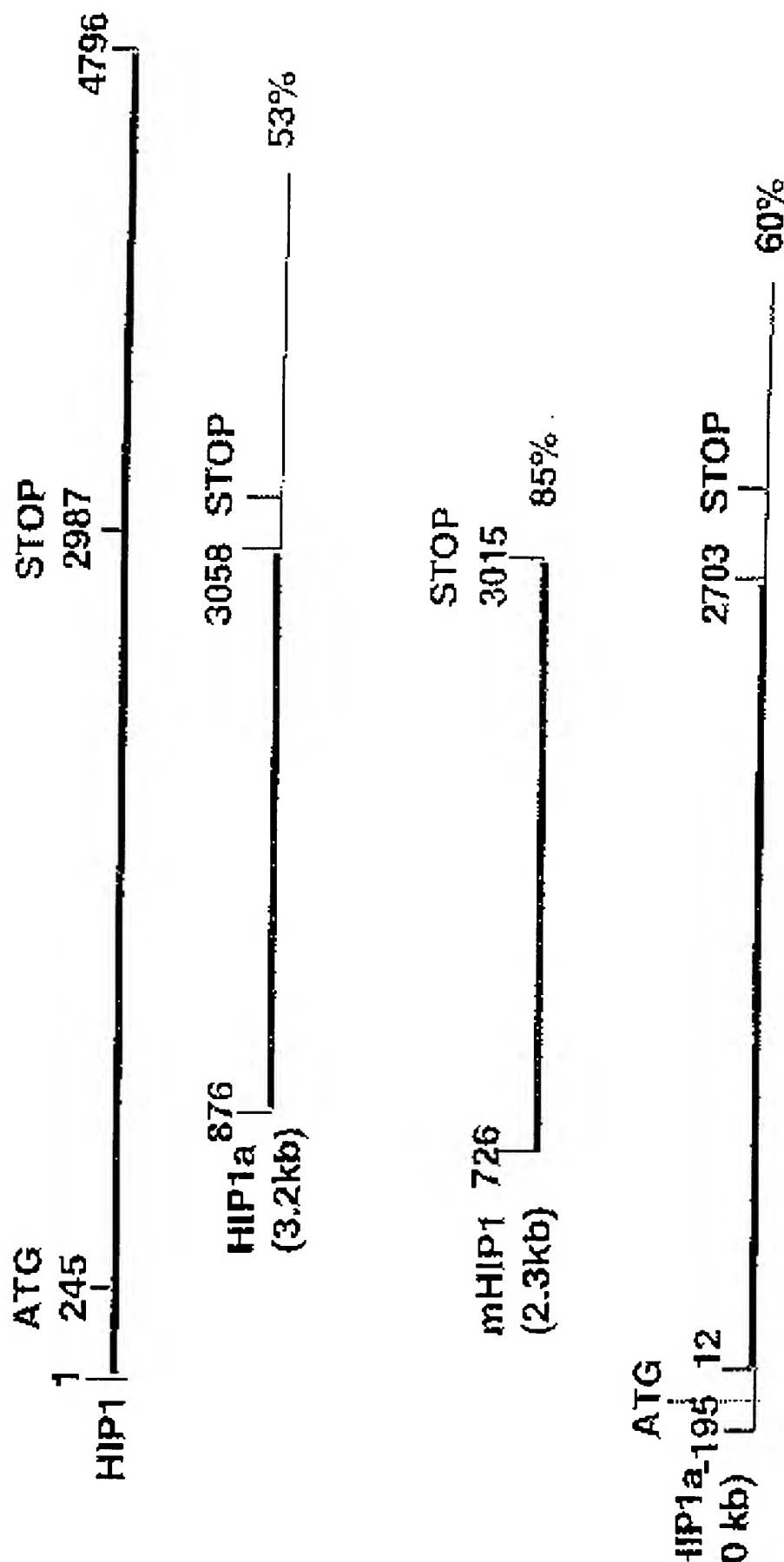


Fig 3
HIP1 Clones: Protein Alignment

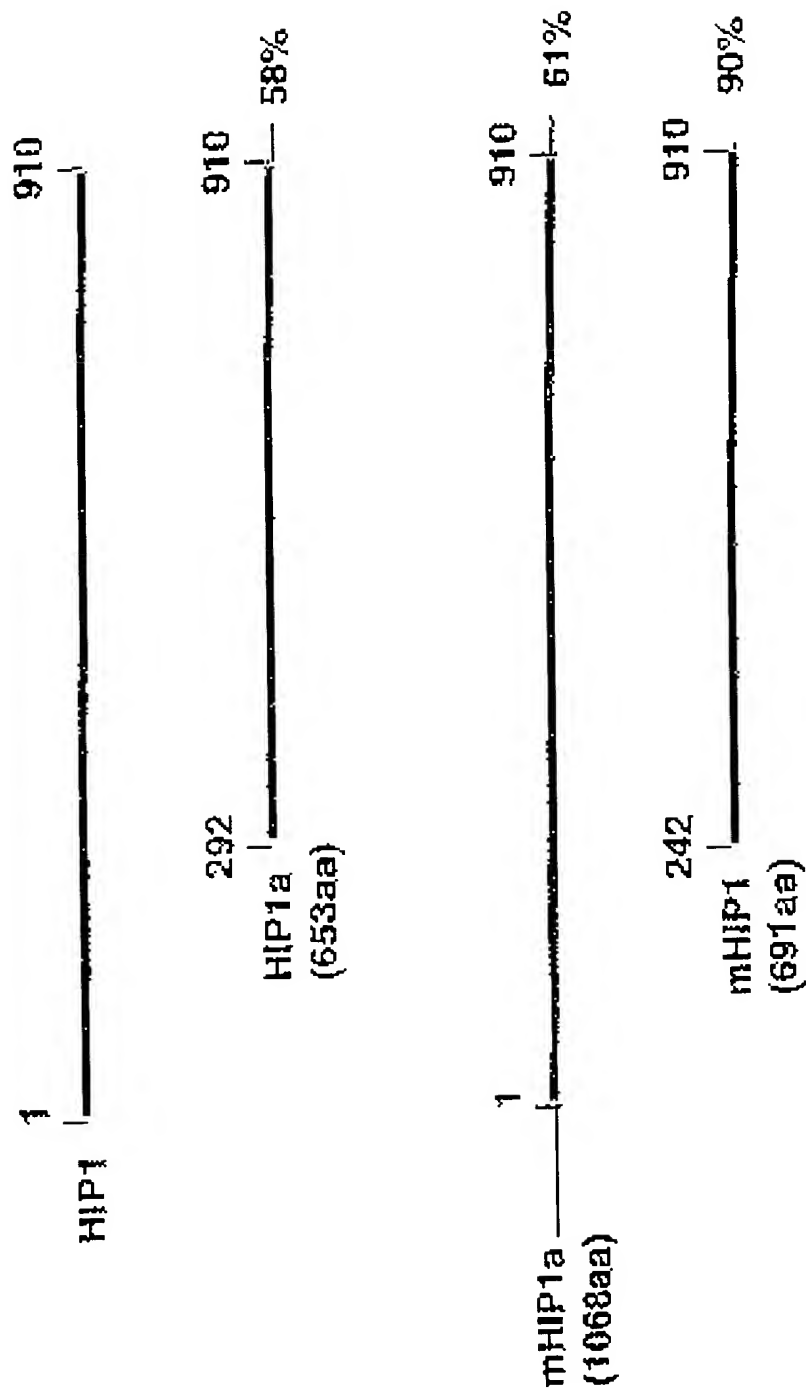


Fig 4

>Usurpin A
SAEVTBQVEENLDTEKKMLLELCRDVAIDVVPFVADLLDILREERKILVCDLRELLYRVEREDILKRLK

>Usurpin B
YRVLMAHIGEDLDKSDVSSLIFLMKTYMGSGKYSKHKSFLDLVVELKKLVLPDQLDILEKCLKNIHRI DLYKYKQK

>Casp-0 A
FSRNLYDIGETQDSEDLASLKELSDYTPQRKDETIKIALMTQPIQEKPMLEESNLSFLKELLFRINRLDLLITYLN

>Casp-8 B
YRWLYQISEEVSAREIRSEFKFLQSELSKCHLDDQHNLLQIFEMKKVILGCKLDILKKVCAQINKSLFKIND

>Casp-10 A
FKKKLL7IDSNLGVDVENIKFLCJGLVFNKKLDRSSASDVFEHLLADLLSESDPFTIABLIYYTSQKKISQHLNC

>Casp-10 B
FRNLLYELSGSTDSRWLKDHTFLKXDDLKTEMTSLSTAFLLKXGXTDEDNLTCLDLCXTVVPKLLRNIEK

>FADD
FLVQLNSVSSSLSSSELTFLKFLCLGRVGRKKLERVQSGLDLFSMLLEQNDLEDSITELASLLASTRRDILLRYDD

>MC159 A
SLPFLRHLLDELDSEDSLLSYGHDAAPGCTTVTQALCSLSQQKSLTLAALVEMLYVLRMDLLKSRNG

>MC159 B
YRKEMVGVGEEDDSSELKALRTFACNMPGLSTALSSESRPVEIQLALENVGLVSPSSVSYLADMLRTIRRLDLCQQLVE

>PB
FRCLMALVNDFLSNKVEEYFLCAPLESHEPSSKKSFRLASLLEDLLELGGDKLTLRHLLTTIGRADIVANLQV

>K3 orf13A
TYEVICEVARKLGTDREVVLFLLVVFLLPQPTLAQILGALRALKEGRITFFLLAECLPRAGBRULLRDLH

>K5 orf13B
YQLTVLHYDGEICARDIRSLIFLSKDTIGSRSTPQVYLFNVYOMENIDILCPTDVAI MSNLSLSRVDIQRQVQT

>HIP1
SELFAQLAEQQHLRQQAADDCETLSAFLNELLARQREDTEKAQASLSSTERKAQANEQRYSLKKEKYSELVQNHADILRKN
AS

>HIP1a
GEYSPQRKQKQKALVDNEQLRHELAQLRQAQLERERSQGLREDAERKASATEARYNKIKKKHSELVIVHAELLRNAD

>mHIP1a
NGLEAZLLEQRKQKQKALVDNEQLRHELAQLNALQLEBARHUGLREDAERKASATEARYSKLKKHSEIINTINELLRKN
AD

>mHIP1
SELEBELAEQQHLGRQAMDUCEFIETFEIDRIKQREDTEKAQASLTELKKAQANEQRYSKLKEKYSELVQNHADILRKN
AE

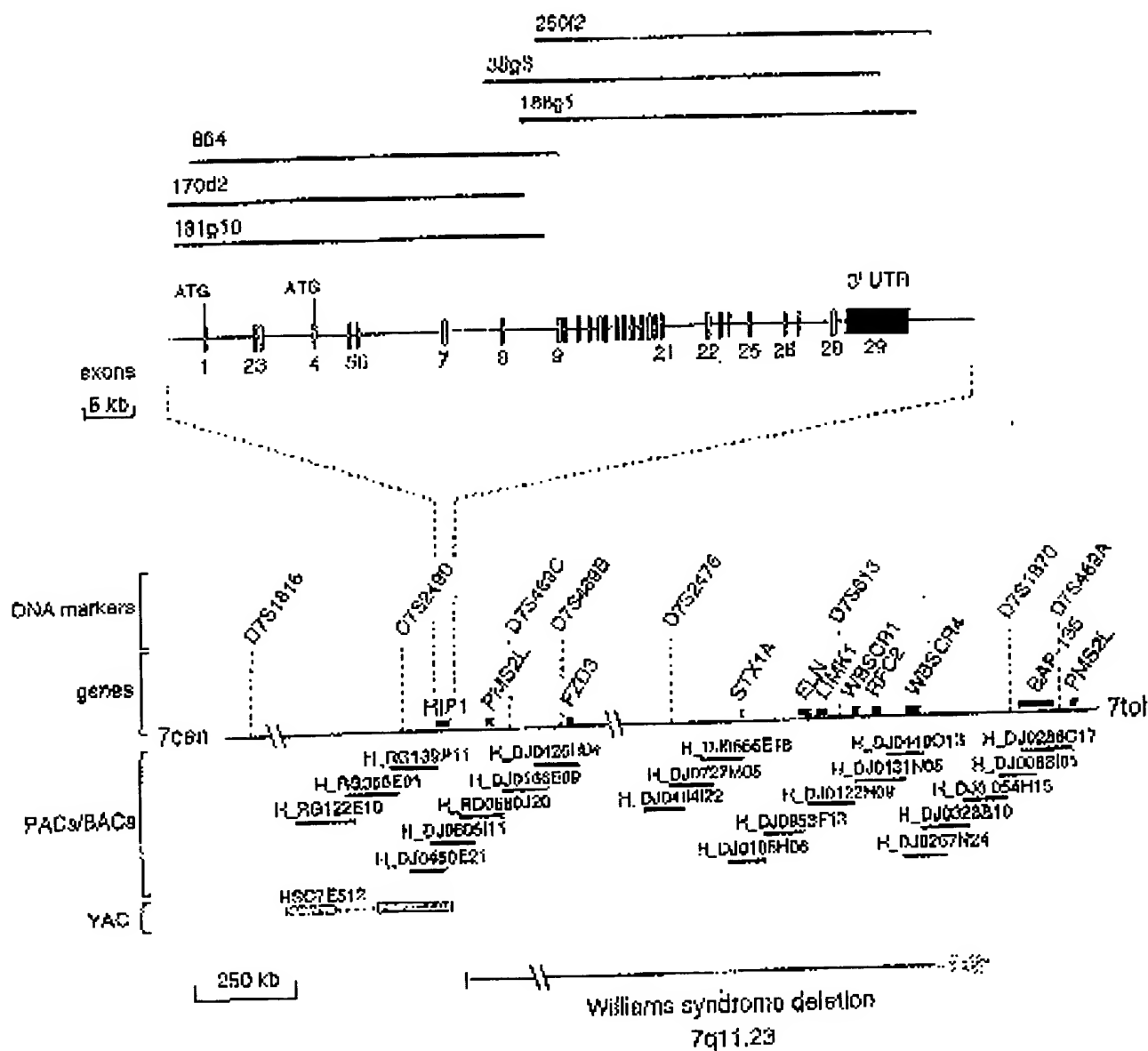


Fig 5

Fig 2

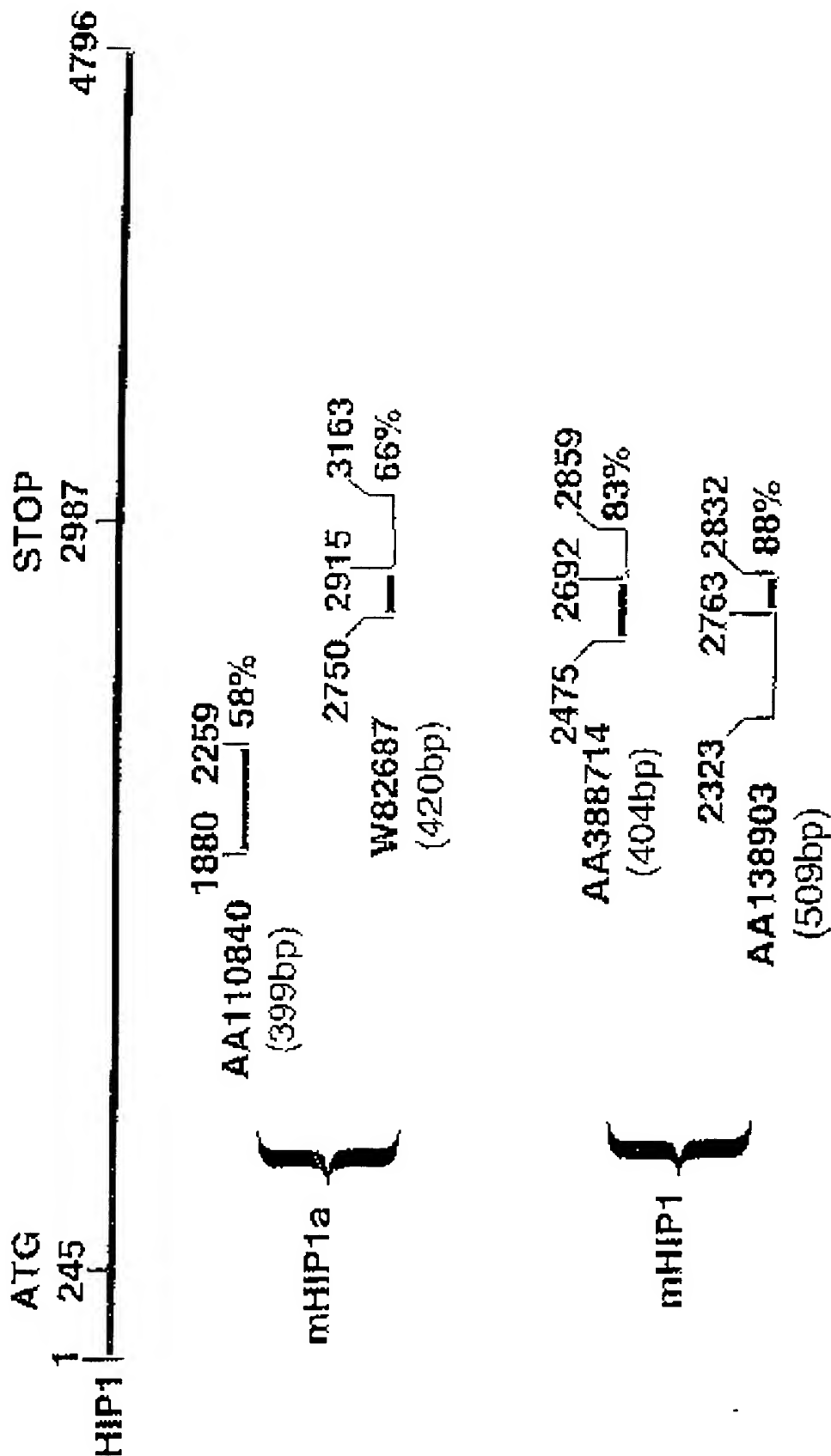
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2	zk370.3	279
1	h1p1	445
2	zk370.3	345
1	h1p1	530
2	zk370.3	429
1	h1p1	517
2	zk370.3	498
1	h1p1	702
2	zk370.3	555
1	h1p1	792
2	zk370.3	661
1	h1p1	882
2	zk370.3	731

FIG 6 (CONT.)

[illegible]

Fig 7

Mouse ESTs



mtdjp1.pains@raph2 Tue Apr 26 11:30:41 1999

Hip1 increases the susceptibility to cell stress

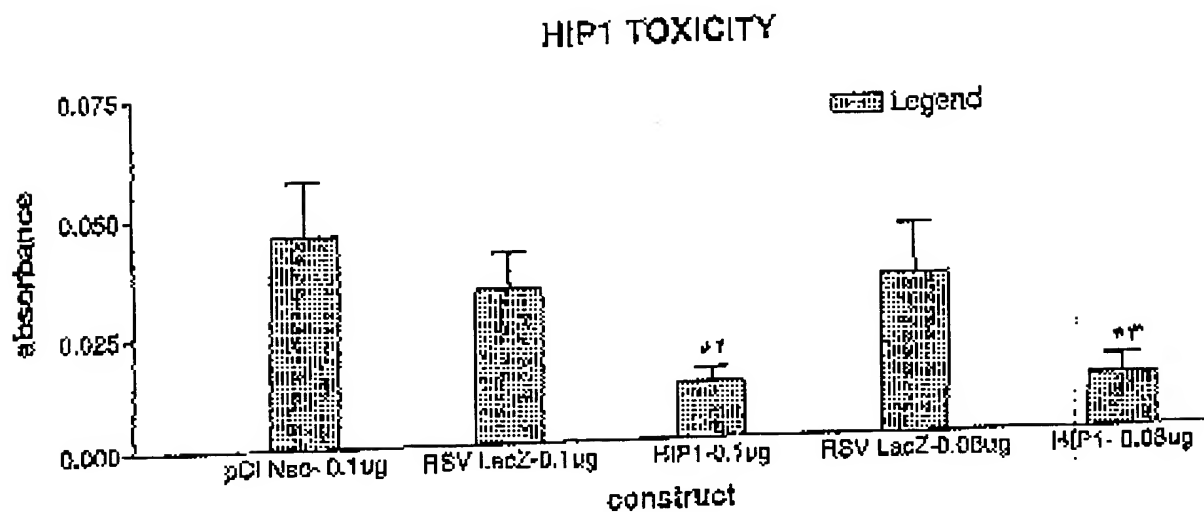


Fig 8

Hip-1 is toxic in the presence of huntingtin

HIP1 transfected into HD1955-15 stable cell line
36 hr post-transfection

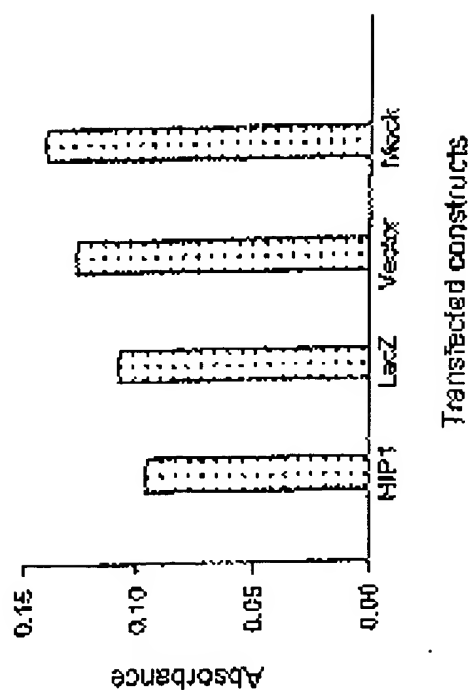


Fig 9A

Hip-1 is toxic in the presence of knockdown

HIP1 transfected into HD1955-128 stable cell line
36 hr post-transfection

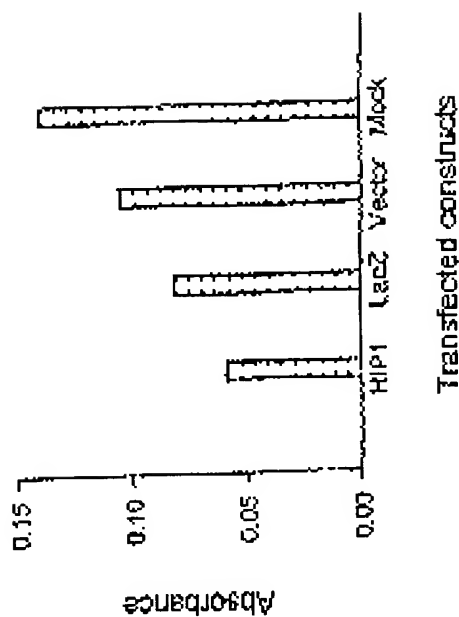
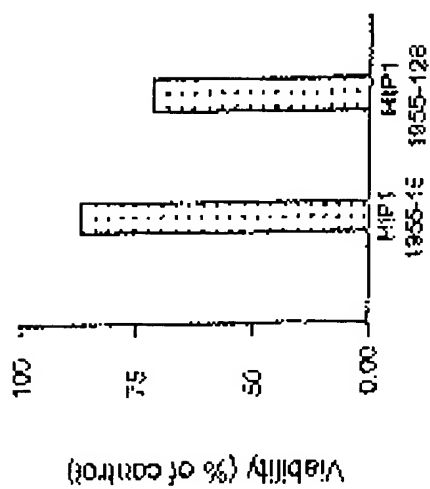


Fig 9B

Polyglutamine-dependence of HIP-1 toxicity



Transfected constructs/cell lines

Fig 9C

SEQUENCE LISTING

(1) GENERAL INFORMATION:

(i) APPLICANT: Kalchman, Michael

Hayden, Michael R.

Hackam, Abigail

Chopra, Vikramjit Singh

Nicholson, Donald W.

Vallaincourt, John P.

Raspor, Dita M.

(ii) TITLE OF INVENTION: Apoptosis Modulators That Interact with the
Huntington's Disease Gene

(iii) NUMBER OF SEQUENCES: 44

(iv) CORRESPONDENCE ADDRESS:

(A) ADDRESSEE: Oppedahl & Larson

(B) STREET: PO Box 5270

(C) CITY: Frisco

(D) STATE: CO

(E) COUNTRY: USA

(F) ZIP: 80443-5270

(v) COMPUTER READABLE FORM:

(A) MEDIUM TYPE: Diskette, 3.50 inch, 1.44 Kb storage

(B) COMPUTER: IBM Compatible

(C) OPERATING SYSTEM: MS DOS 5.0

(D) SOFTWARE: WordPerfect

(vi) CURRENT APPLICATION DATA:

(A) APPLICATION NUMBER:

(B) FILING DATE:

(C) CLASSIFICATION:

(viii) ATTORNEY/AGENT INFORMATION:

(A) NAME: Larson, Marina T.

(B) REGISTRATION NUMBER: 32038

(C) REFERENCE/DOCKET NUMBER: UBC.P-013US2

(ix) TELECOMMUNICATION INFORMATION:

(A) TELEPHONE: (970) 668-2050

(B) TELEFAX: (970) 668-2052

(2) INFORMATION FOR SEQ ID NO: 1:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1164

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: cDNA for Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

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GCCTCAGCCC TGTGAGAAAC TATCAGCCCT GTGGTGGTGA TCCCTGCAGA   200
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TTTGGCAGTT CATCCAGCAG TGATCCCTTC AATTTCACAA GTCAAAATGG   350
TGTGAACAAG GATGAGAAGC ACCAATTAAAT TGAGCGAATA TACAGAGAGA   400
TCAGTGGATT GAAGGCACAC CTAGAAAACA TGAAGACTUA GAGCCAGCGG   450
GTTGTGCTGC AGCTGAAGGG CCACGTCAGC GAGCTGGAAG CAGATCTGGC   500
CGAGCAGCAG CACCTGCGGC AGCAGGCGGC CGACGACTGT GAATTCCTGC   550
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CAGCGGAGCC TGTCTGAGAT AGAAAGGAAA GCTCAAGCCA ATGAACAGCG   650
ATATAGCAAG CTAAAGGAGA AGTACAGCGA CCTCCTTCAG AACCACTCTG   700
ACCTGCTGCG GAAGAATGCA CAGGTGAUUA AACAGGTGTC CATGCCCAGA   750
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GGAGCGCATC AGTGACCAGG GCCAGCGGAA GACTCAAGAA CAGCTGGAAG   850
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CTTC TGCAAG GCAGCCTGGA AACTTCTGCC CAGTCAGAAG CAAACTGGGC   950
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CAGCTCATAG GGAGGAGGAA TTATCTGCTC TTCGGAAAGA ACTGCAGGAC 1050
ACTCAGCTCA AACTGCCCAAG CACAGAGGAA TCTATGTGCC AGCTTGCCAA 1100
AGACCAACGA AAAATGCTTC TGGTGGGGTC CAGGAAGGCT GCGGACCAGG 1150
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(2) INFORMATION FOR SEQ ID NO:2:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 386

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

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Phe Thr Lys Leu Lys Asp Leu Phe Tyr Arg Ser Ser Asn Leu Gln
                20             25             30

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Tyr Phe Lys Arg Val Ile Gln Ile Pro Gln Leu Pro Glu Asn Pro

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Val Val Val Ile Pro Ala Glu Ala Ser Ser Pro Asp Ser Glu Pro	65	70	75
Val Leu Glu Lys Asp Asp Leu Met Asp Met Asp Ala Ser Gln Gln	80	85	90
Asn Leu Phe Asp Asn Lys Phe Asp Asp Phe Gly Ser Ser Ser Ser	95	100	105
Ser Asp Pro Phe Asn Phe Asn Ser Gln Asn Gly Val Asn Lys Asp	110	115	120
Glu Lys Asp His Leu Ile Glu Arg Leu Tyr Arg Glu Ile Ser Gly	125	130	135
Leu Lys Ala Gln Leu Glu Asn Met Lys Thr Glu Ser Gln Arg Val	140	145	150
Val Leu Gln Leu Lys Gly His Val Ser Glu Leu Glu Ala Asp Leu	155	160	165
Ala Glu Gln Gln His Leu Arg Gln Gln Ala Ala Asp Asp Cys Glu	170	175	180
Phe Leu Arg Ala Glu Leu Asp Glu Leu Arg Gln Arg Glu Asp Thr	185	190	195
Glu Lys Ala Gln Arg Ser Leu Ser Glu Ile Glu Arg Lys Ala Gln	200	205	210
Ala Asn Glu Gln Arg Tyr Ser Lys Leu Lys Glu Lys Tyr Ser Glu	215	220	225
Leu Val Gln Asn His Ala Asp Leu Leu Arg Lys Asn Ala Glu Val	230	235	240
Thr Lys Gln Val Ser Met Ala Arg Gln Ala Gln Val Asp Leu Glu	245	250	255
Arg Glu Lys Lys Glu Leu Glu Asp Ser Leu Glu Arg Ile Ser Asp	260	265	270
Gln Gly Gln Arg Lys Thr Gln Gln Gln Leu Glu Val Leu Glu Ser	275	280	285
Leu Lys Gln Glu Leu Gly Thr Ser Gln Arg Glu Leu Gln Val Leu			

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Gln Gly Ser Leu	Glu Thr Ser Ala Gln	Ser Gln Ala Asn Trp	Ala		
	305		310		315
Ala Glu Phe Ala	Glu Leu Glu Lys Glu	Arg Asp Ser Leu Val	Ser		
	320		325		330
Gly Ala Ala His	Arg Glu Glu Glu Leu	Ser Ala Leu Arg Lys	Glu		
	335		340		345
Leu Gln Asp Thr	Gln Leu Lys Leu Ala	Ser Thr Glu Glu Ser	Met		
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Cys Gln Leu Ala	Lys Asp Gln Arg Lys	Met Leu Leu Val Gly	Ser		
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Arg Lys Ala Ala	Glu Gln Val Ile Gln	Asp Ala			
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(2) INFORMATION FOR SEQ ID NO:3:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 4796

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: cDNA for Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:

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TTGCACCCAC	CATGAGAAAG	GGGCAACAGA	CCCTNTGTIN	TGTINTCAAC	200
CGCCTGCTTC	TGTTTTAGCA	ACGCAGTGIT	TTGGTGC AAC	TTGTCCCATG	250
TGTTCCACAA	ANTCTTCCGA	GATCGACACU	CGAACGTCCU	GAAGGACTTT	300
GTGAGATACA	GAAATGAATT	GAGTGACATG	AGCAGGATGT	GGGGCCACCT	350
GAGCGAGGGG	TATGGCCAGC	TGTGCAGCAT	CTACCTGAAA	CTGCTAAGAA	400
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ATGAGTGACC	GCCAGCTGGA	CGAGGCTGGA	GAAAGTGACG	TGAACAACIT	500
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TCTTCCAAAC	AGTATTCAAC	TCCCTGGACA	TGTCCCGCTC	TGTGTCCGTG	600
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CCTGCCTCCC	AGCTGACACC	CTGCAAGGCC	ACCEGCACCG	CTTCATGGAG	750

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ATGACATCTT	TGGCAGTTCA	TTCAGCAGTG	ATCCCTTCAA	TTCACACAGT	1050
CAAAATGGTG	TGAACAAGGA	TGAGAAGGAC	CACCTTAAATC	AGCGACTATA	1100
CAGACAGATC	AGTGGATTGA	AGGCACAGCT	ACAAAACATG	AAGACTGAGA	1150
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CCTAGCCCAG	CTGCAGCAGG	CCTCTCGGGG	AGTGAAACCAG	GCCACTGCCG	2800
GCGTTGTGGC	CTCAACCAAT	TCCGGCAAA	CACACATCGA	AGACACAGAC	2850
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GGTAGCTCCA	GCGATGGTGC	TGCCCAGGCC	TCTCGGTGCT	CCATCTCCGC	3850
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GATCGTTCCA	CTTCTACGC	AATTGACAAA	CCCGGAAGAT	CAGATGCAT	4300
TGCTCCCATC	AGGGAAGAAC	CCTATACTTG	GTTTGCTACC	CTTAGTAAAT	4350
ATTACTAACC	TCCCTTAAGC	AGCAACAGCC	TACAAAGAGA	TGCTTCCAGC	4400
AAATCAGAACT	TCAGGTGTGA	CTCTAGCAAA	GCTCATCTTT	CTGCCCCGCT	4450
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ATGAGAAAGC	ACAGAGCCAG	CGCCTCCAAC	TCCTTTCAGC	CACATGCCCT	4600
AGGCTCTCGC	TGCCCTGTGG	ACAGGATGAG	GACAGAGGGC	ACATGAACAG	4650
CTTGCCAGGG	ATGGGCAGCC	CAACAGCACT	TTTCCTCTTC	TAGATGGACC	4700
CCAGCATTTA	AGTGACCTTC	TGATCTTGGG	AAAACAGCGT	CTTCCTTCTT	4750
TATCTATAGC	AATCATTTGG	TGGTAGCCAT	CAAGCACTTC	GGAAAT	4795

(2) INFORMATION FOR SEQ ID NO: 4:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 924

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:

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				20				25						30

Thr	Lys	Asn	Pro	Arg	Phe	Pro	Gly	Asn	Leu	Gln	Met	Ser	Asp	Arg			
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Gln	Leu	Asp	Glu	Ala	Gly	Glu	Ser	Asp	Val	Asn	Asn	Phe	Phe	Gln			
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Phe	Gln	Thr	Val	Phe	Asn	Ser	Leu	Asp	Met	Ser	Arg	Ser	Val	Ser			
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Val	Thr	Ala	Ala	Gly	Gln	Cys	Arg	Leu	Ala	Pro	Leu	Ile	Gln	Val			
				95					100					105			
Ile	Leu	Asp	Cys	Ser	His	Leu	Tyr	Asp	Tyr	Thr	Val	Lys	Leu	Leu			
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Phe	Lys	Leu	His	Ser	Cys	Leu	Pro	Ala	Asp	Thr	Leu	Gln	Gly	His			
				125					130					135			
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				140					145					150			
Tyr	Arg	Ser	Ser	Asn	Leu	Gln	Tyr	Phe	Lys	Arg	Leu	Ile	Gln	Ile			
				155					160					165			
Pro	Gln	Leu	Pro	Glu	Asn	Pro	Pro	Asn	Phe	Leu	Arg	Ala	Ser	Ala			
				170					175					180			
Leu	Ser	Glu	His	Ile	Ser	Pro	Val	Val	Val	Ile	Pro	Ala	Glu	Ala			
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Ser	Ser	Pro	Asp	Ser	Glu	Pro	Val	Leu	Glu	Lys	Asp	Asp	Leu	Met			
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Asp	Met	Asp	Ala	Ser	Gln	Gln	Asn	Leu	Phe	Asp	Asn	Lys	Phe	Asp			
				215					220					225			
Asp	Ile	Phe	Gly	Ser	Ser	Phe	Ser	Ser	Asp	Pro	Phe	Asn	Phe	Asn			
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Ser	Gln	Asn	Gly	Val	Asn	Lys	Asp	Glu	Lys	Asp	His	Leu	Ile	Glu			
				245					250					255			
Arg	Leu	Tyr	Arg	Glu	Ile	Ser	Gly	Leu	Lys	Ala	Gln	Leu	Glu	Asn			
				260					265					270			
Met	Lys	Thr	Glu	Ser	Gln	Arg	Val	Val	Leu	Gln	Leu	Lys	Gly	His			
				275					280					285			

Val	Ser	Glu	Leu	Glu	Ala	Asp	Leu	Ala	Glu	Gln	Gln	His	Leu	Arg	290	295	300
Gln	Gln	Ala	Ala	Asp	Asp	Cys	Clu	Phe	Leu	Arg	Ala	Glu	Leu	Asp	305	310	315
Glu	Leu	Arg	Arg	Gln	Arg	Glu	Asp	Thr	Glu	Lys	Ala	Gln	Arg	Ser	320	325	330
Leu	Ser	Glu	Ile	Glu	Arg	Lys	Ala	Gln	Ala	Asn	Glu	Gln	Arg	Tyr	335	340	345
Ser	Lys	Leu	Lys	Glu	Lys	Tyr	Ser	Glu	Leu	Val	Gln	Asn	His	Ala	350	355	360
Asp	Leu	Leu	Arg	Lys	Asn	Ala	Glu	Val	Thr	Lys	Gln	Val	Ser	Met	365	370	375
Ala	Arg	Gln	Ala	Gln	Val	Asp	Leu	Clu	Arg	Glu	Lys	Lys	Glu	Leu	380	385	390
Glu	Asp	Ser	Leu	Glu	Arg	Ile	Ser	Asp	Gln	Gly	Gln	Arg	Lys	Thr	395	400	405
Gln	Clu	Gln	Leu	Glu	Val	Leu	Glu	Ser	Leu	Lys	Gln	Glu	Leu	Gly	410	415	420
Thr	Ser	Gln	Arg	Glu	Leu	Gln	Val	Leu	Gln	Gly	Ser	Leu	Glu	Thr	425	430	435
Ser	Ala	Gln	Ser	Glu	Ala	Asn	Trp	Ala	Ala	Glu	Phe	Ala	Glu	Leu	440	445	450
Glu	Lys	Glu	Arg	Asp	Ser	Leu	Val	Ser	Gly	Ala	Ala	His	Arg	Glu	455	460	465
Glu	Glu	Leu	Ser	Ala	Leu	Arg	Lys	Glu	Leu	Gln	Asp	Thr	Gln	Leu	470	475	480
Lys	Leu	Ala	Ser	Thr	Glu	Glu	Ser	Met	Cys	Gln	Leu	Ala	Lys	Asp	485	490	495
Gln	Arg	Lys	Met	Leu	Leu	Val	Gly	Ser	Arg	Lys	Ala	Ala	Glu	Gln	500	505	510
Val	Ile	Gln	Asp	Ala	Leu	Asn	Gln	Leu	Glu	Glu	Pro	Pro	Leu	Ile	515	520	525
Ser	Cys	Ala	Gly	Ser	Ala	Asp	His	Leu	Leu	Ser	Thr	Val	Thr	Ser	530	535	540

Ile	Ser	Ser	Cys	Ile	Glu	Gln	Leu	Glu	Lys	Ser	Trp	Ser	Gln	Tyr
				545					550					555
Leu	Ala	Cys	Pro	Glu	Asp	Ile	Ser	Gly	Leu	Leu	His	Ser	Ile	Thr
				560					565					570
Leu	Leu	Ala	His	Leu	Thr	Ser	Asp	Ala	Ile	Ala	His	Gly	Ala	Thr
				575					580					585
Thr	Cys	Leu	Arg	Ala	Pro	Pro	Glu	Pro	Ala	Asp	Ser	Leu	Thr	Glu
				590					595					600
Ala	Cys	Lys	Gln	Tyr	Gly	Arg	Glu	Thr	Leu	Ala	Tyr	Leu	Ala	Ser
				605					610					615
Leu	Glu	Glu	Glu	Gly	Ser	Leu	Glu	Asn	Ala	Asp	Ser	Thr	Ala	Met
				620					625					630
Arg	Asn	Cys	Leu	Ser	Lys	Ile	Lys	Ala	Ile	Gly	Glu	Glu	Leu	Leu
				635					640					645
Pro	Arg	Gly	Leu	Asp	Ile	Lys	Gln	Glu	Glu	Leu	Gly	Asp	Leu	Val
				650					655					660
Asp	Lys	Glu	Met	Ala	Ala	Thr	Ser	Ala	Ala	Ile	Glu	Thr	Cys	Thr
				665					670					675
Ala	Arg	Ile	Glu	Glu	Met	Leu	Ser	Lys	Ser	Arg	Ala	Gly	Asp	Thr
				680					685					690
Gly	Val	Lys	Leu	Glu	Val	Asn	Glu	Arg	Ile	Leu	Arg	Cys	Cys	Thr
				695					700					705
Ser	Leu	Met	Gln	Ala	Ile	Gln	Val	Leu	Ile	Val	Ala	Ser	Lys	Asp
				710					715					720
Leu	Gln	Arg	Glu	Ile	Val	Glu	Ser	Gly	Arg	Gly	Thr	Ala	Ser	Pro
				725					730					735
Lys	Glu	Phe	Tyr	Ala	Lys	Asn	Ser	Arg	Trp	Thr	Glu	Gly	Leu	Ile
				740					745					750
Ser	Ala	Ser	Lys	Ala	Val	Gly	Trp	Gly	Ala	Thr	Val	Met	Val	Asp
				765					770					775
Ala	Ala	Asp	Leu	Val	Val	Gln	Gly	Arg	Gly	Lys	Phe	Glu	Glu	Leu
				780					785					790
Met	Val	Cys	Ser	His	Glu	Ile	Ala	Ala	Ser	Thr	Ala	Gln	Leu	Val
				795					800					805

Ala	Ala	Ser	Lys	Val	Lys	Ala	Asp	Lys	Asp	Ser	Pro	Asn	Leu	Ala	
				810					815					820	
Gln	Leu	Gln	Gln	Ala	Ser	Arg	Gly	Val	Asn	Gln	Ala	Thr	Ala	Gly	
				825					830					835	
Val	Val	Ala	Ser	Thr	Ile	Ser	Gly	Lys	Ser	Gln	Ile	Glu	Glu	Thr	
				840					845					850	
Asp	Asn	Met	Asp	Phe	Ser	Ser	Met	Thr	Leu	Thr	Gln	Ile	Lys	Arg	
				855					860					865	
Gln	Glu	Met	Asp	Ser	Gln	Val	Arg	Val	Leu	Glu	Leu	Glu	Asn	Glu	
				870					875					880	
Leu	Gln	Lys	Glu	Arg	Gln	Lys	Leu	Gly	Glu	Leu	Arg	Lys	Lys	His	
				885					890					895	
Tyr	Glu	Leu	Ala	Gly	Val	Ala	Glu	Gly	Trp	Glu	Glu	Gly	Thr	Glu	
				900					905					910	
Ala	Ser	Pro	Pro	Thr	Leu	Gln	Glu	Val	Val	Thr	Glu	Lys	Glu		
				915					920				924		

(2) INFORMATION FOR SEQ ID NO: 5

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1090

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: Huntinglin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:

Met	Leu	Leu	Cys	Gln	Gly	Ser	Glu	Trp	Arg	Arg	Asp	Gln	Gln	Leu	
				5					10					15	
Gly	Thr	Ala	Asn	Ala	Arg	Gln	Trp	Cys	Pro	Leu	Pro	Gln	Asp	Ala	
				20					25					30	
Gln	Pro	Ala	Gly	Ser	Trp	Glu	Arg	Cys	Pro	Pro	Leu	Pro	Pro	Ala	
				35					40					45	
Gly	Arg	Leu	Gln	Gly	Thr	Asp	His	Pro	Trp	Gly	Trp	Gly	Arg	Leu	
				50					55					60	

Ala	Gly	Gly	Gly	Glu	Arg	Gly	Gly	Leu	Trp	Glu	Gly	Leu	Ser	His	65	70	75
Ser	Gln	Arg	Leu	Ile	His	Leu	Ile	Leu	Leu	Ser	Leu	Pro	Leu	Leu	80	85	90
Val	Phe	Gln	Thr	Val	Ser	Ile	Asn	Lys	Ala	Ile	Asn	Thr	Gln	Glu	95	100	105
Val	Ala	Val	Lys	Glu	Lys	His	Ala	Arg	Thr	Cys	Ile	Leu	Gly	Thr	110	115	120
His	His	Glu	Lys	Gly	Ala	Gln	Thr	Phe	Trp	Ser	Val	Val	Asn	Arg	125	130	135
Leu	Pro	Leu	Ser	Ser	Asn	Ala	Val	Leu	Cys	Trp	Lys	Phe	Cys	His	140	145	150
Val	Phe	His	Lys	Leu	Leu	Arg	Asp	Gly	His	Pro	Asn	Val	Leu	Lys	155	160	165
Asp	Ser	Leu	Arg	Tyr	Arg	Asn	Glu	Leu	Ser	Asp	Met	Ser	Arg	Met	170	175	180
Trp	Gly	His	Leu	Ser	Glu	Gly	Tyr	Gly	Gln	Leu	Cys	Ser	Ile	Tyr	185	190	195
Leu	Lys	Leu	Leu	Arg	Thr	Lys	Met	Glu	Tyr	His	Thr	Lys	Asn	Pro	200	205	210
Arg	Phe	Pro	Gly	Asn	Leu	Gln	Met	Ser	Asp	Arg	Gln	Leu	Asp	Glu	215	220	225
Ala	Gly	Glu	Ser	Asp	Val	Asn	Asn	Phe	Phe	Gln	Leu	Thr	Val	Glu	230	235	240
Met	Phe	Asp	Tyr	Leu	Glu	Cys	Glu	Leu	Asn	Leu	Phe	Gln	Thr	Val	245	250	255
Phe	Asn	Ser	Leu	Asp	Met	Ser	Arg	Ser	Val	Ser	Val	Thr	Ala	Ala	260	265	270
Gly	Gln	Cys	Arg	Leu	Ala	Pro	Leu	Ile	Gln	Val	Ile	Leu	Asp	Cys	275	280	285
Ser	His	Leu	Tyr	Asp	Tyr	Thr	Val	Lys	Leu	Leu	Phe	Lys	Leu	His	290	295	300
Ser	Cys	Leu	Pro	Ala	Asp	Thr	Leu	Gln	Gly	His	Arg	Asp	Arg	Phe	305	310	315

Met	Glu	Gln	Phe	Thr	Lys	Leu	Lys	Asp	Leu	Phe	Tyr	Arg	Ser	Ser	320	325	330
Asn	Leu	Gln	Tyr	Phe	Lys	Arg	Leu	Ile	Gln	Ile	Pro	Gln	Leu	Pro	335	340	345
Glu	Asn	Pro	Pro	Asn	Phe	Leu	Arg	Ala	Ser	Ala	Leu	Ser	Glu	His	350	355	360
Ile	Ser	Pro	Val	Val	Val	Ile	Pro	Ala	Glu	Ala	Ser	Ser	Pro	Asp	365	370	375
Ser	Glu	Pro	Val	Leu	Glu	Lys	Asp	Asp	Leu	Met	Asp	Met	Asp	Ala	380	385	390
Ser	Gln	Gln	Asn	Leu	Phe	Asp	Asn	Lys	Phe	Asp	Asp	Ile	Phe	Gly	395	400	405
Ser	Ser	Phe	Ser	Ser	Asp	Pro	Phe	Asn	Phe	Asn	Ser	Gln	Asn	Gly	410	415	420
Val	Asn	Lys	Asp	Glu	Lys	Asp	His	Leu	Ile	Glu	Arg	Leu	Tyr	Arg	425	430	435
Glu	Ile	Ser	Gly	Leu	Lys	Ala	Gln	Leu	Glu	Asn	Met	Lys	Thr	Glu	440	445	450
Ser	Gln	Arg	Val	Val	Leu	Glu	Leu	Lys	Gly	His	Val	Ser	Glu	Leu	455	460	465
Glu	Ala	Asp	Leu	Ala	Glu	Gln	Gln	His	Leu	Arg	Gln	Gln	Ala	Ala	470	475	480
Asp	Asp	Cys	Glu	Phe	Leu	Arg	Ala	Glu	Leu	Asp	Glu	Leu	Arg	Arg	485	490	495
Gln	Arg	Glu	Asp	Thr	Glu	Lys	Ala	Gln	Arg	Ser	Leu	Ser	Glu	Ile	500	505	510
Glu	Arg	Lys	Ala	Gln	Ala	Asn	Glu	Gln	Arg	Tyr	Ser	Lys	Leu	Lys	515	520	525
Glu	Lys	Tyr	Ser	Glu	Leu	Val	Gln	Asn	His	Ala	Asp	Leu	Leu	Arg	530	535	540
Lys	Asn	Ala	Glu	Val	Thr	Lys	Gln	Val	Ser	Met	Ala	Arg	Gln	Ala	545	550	555
Gln	Val	Asp	Leu	Glu	Arg	Glu	Lys	Lys	Glu	Leu	Glu	Asp	Ser	Leu	560	565	570

Glu Arg Ile Ser	Asp Gln Gly Gln Arg	Lys Thr Gln Glu Gln Leu	575	588	585
Glu Val Leu Glu	Ser Leu Lys Gln Glu	Leu Ala Thr Ser Gln Arg	590	595	600
Glu Leu Gln Val	Leu Gln Gly Ser Leu	Glu Thr Ser Ala Gln Ser	605	610	615
Glu Ala Asn Trp	Ala Ala Glu Phe Ala	Glu Leu Glu Lys Glu Arg	620	625	630
Asp Ser Leu Val	Ser Gly Ala Ala His	Arg Glu Glu Glu Leu Ser	635	640	645
Ala Leu Arg Lys	Glu Leu Gln Asp Thr	Gln Leu Lys Leu Ala Ser	650	655	660
Thr Glu Glu Ser	Met Cys Gln Leu Ala	Lys Asp Gln Arg Lys Met	665	670	675
Leu Leu Val Gly	Ser Arg Lys Ala Ala	Glu Gln Val Ile Gln Asp	680	685	690
Ala Leu Asn Gln	Leu Glu Glu Pro Pro	Leu Ile Ser Cys Ala Gly	695	700	705
Ser Ala Asp His	Leu Leu Ser Thr Val	Thr Ser Ile Ser Ser Cys	710	715	720
Ile Glu Gln Leu	Glu Lys Ser Trp Ser	Gln Tyr Leu Ala Cys Pro	725	730	735
Glu Asp Ile Ser	Gly Leu Leu His Ser	Ile Thr Leu Leu Ala His	740	745	750
Leu Thr Ser Asp	Ala Ile Ala His Gly	Ala Thr Thr Cys Leu Arg	755	760	765
Ala Pro Pro Glu	Pro Ala Asp Ser Leu	Thr Glu Ala Cys Lys Gln	770	775	780
Tyr Gly Arg Glu	Thr Leu Ala Tyr Leu	Ala Ser Leu Glu Glu Glu	785	790	795
Gly Ser Leu Glu	Asn Ala Asp Ser Thr	Ala Met Arg Asn Cys Leu	800	805	810
Ser Lys Ile Lys	Ala Ile Gly Glu Glu	Leu Leu Pro Arg Gly Leu	815	820	825

Asp	Ile	Lys	Gln	Glu	Gln	Leu	Gly	Asp	Leu	Val	Asp	Lys	Glu	Met	830	835	840
Ala	Ala	Thr	Ser	Ala	Ala	Ile	Glu	Thr	Ala	Thr	Ala	Arg	Ile	Glu	845	850	855
Glu	Met	Leu	Ser	Lys	Ser	Arg	Ala	Gly	Asp	Thr	Gly	Val	Lys	Leu	860	865	870
Glu	Val	Asn	Glu	Arg	Ile	Leu	Gly	Cys	Cys	Thr	Ser	Leu	Met	Gln	875	888	885
Ala	Ile	Gln	Val	Leu	Ile	Val	Ala	Ser	Lys	Asp	Leu	Gln	Arg	Glu	890	895	900
Ile	Val	Glu	Ser	Gly	Arg	Gly	Thr	Ala	Ser	Pro	Lys	Glu	Phe	Tyr	905	910	915
Ala	Lys	Asn	Ser	Arg	Trp	Thr	Glu	Gly	Leu	Ile	Ser	Ala	Ser	Lys	920	925	930
Ala	Val	Gly	Trp	Gly	Ala	Thr	Val	Met	Val	Asp	Ala	Ala	Asp	Leu	935	940	945
Val	Val	Gln	Gly	Arg	Gly	Lys	Phe	Glu	Glu	Leu	Met	Val	Cys	Ser	950	955	960
His	Glu	Ile	Ala	Ala	Ser	Thr	Ala	Gln	Leu	Val	Ala	Ala	Ser	Lys	965	970	975
Val	Lys	Ala	Asp	Lys	Asp	Ser	Pro	Asn	Leu	Ala	Gln	Leu	Gln	Gln	980	985	990
Ala	Ser	Arg	Gly	Val	Asn	Gln	Ala	Thr	Ala	Gly	Val	Val	Ala	Ser	995	1000	1005
Thr	Ile	Ser	Gly	Lys	Ser	Gln	Ile	Glu	Glu	Thr	Asp	Asn	Met	Asp	1010	1015	1020
Phe	Ser	Ser	Met	Thr	Leu	Thr	Gln	Ile	Lys	Arg	Gln	Glu	Met	Asp	1025	1030	1035
Ser	Gln	Val	Arg	Val	Leu	Glu	Leu	Glu	Asn	Glu	Leu	Gln	Lys	Glu	1040	1045	1050
Arg	Gln	Lys	Leu	Gly	Glu	Leu	Arg	Lys	Lys	His	Tyr	Glu	Leu	Ala	1055	1060	1065
Gly	Val	Ala	Glu	Gly	Trp	Glu	Glu	Gly	Thr	Glu	Ala	Ser	Pro	Pro	1070	1075	1080

Thr Leu Gln Glu Val Val Thr Glu Lys Glu
1085 1090

(2) INFORMATION FOR SEQ ID NO:6:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 3301

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: cDNA for Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:

CGGTGAGCTC	GAGGAGCAGC	GGAAGCAGAA	GCAGAAGGCC	CTGGTGGATA	50
ATGAGCAGCT	CCCCCAGGAG	CTGGCCCAGC	TGAGGGCTGC	CCAGCTGGAG	100
CGCGAGCGGA	GCCAGGGCCT	GCCTCAGGAG	GCTCAGAGGA	AGGCCAGTGC	150
CACGGAGGCG	CGCTACAACA	AGCTGAAGGA	AAACCCACAGT	GAGCTCGTCC	200
ATGTGCACGC	GGAGCTGCTC	AGAAAGAAAC	CGGACACAGC	CAAGCAGCTG	250
ACGGTGACGC	AGCAAGCCCA	GGAGGAGGTG	GCGCGGGTGA	AGCAGCAGCT	300
GGCCTTCCAC	G'GGAGCAGG	TGAAGCGGGA	GTCGGAGTTG	AAGCTAGAGC	350
AGAAGAGCGA	CCAGCAGGAG	AAGCTCAAGA	GGGAGCTGGA	GGCCAAGGCC	400
GGAGAGCTGG	CCCCCGCGCA	GGAGGCCCTG	AGCCACACAG	AGCAGAGCAA	450
GTCGGAGCTG	AGCTCACGGC	TGGACACACT	GAGTGCGGAG	AAGGATGCTC	500
TGAGTGGAGC	TGTGCGGCAG	CGGGAGGCAG	ACCTGCTGGC	GGCGCAGAGC	550
CTGGTSCGCG	AGACAGAGGC	GGCGCTGAGC	CEGLAGCAGC	AGCGCAGCTC	600
CCAGGAGCAG	GGCGAGTTGC	AGGGCCGGCT	GCCAGAGAGG	GAGTCTCAGG	650
AGCAGGGGCT	CCCGCAGAGG	CTGCTGGACG	AGCAGTTCCG	AGTGTTCGGG	700
GGCGCTGCTG	CCGAGGCCCG	CGGCATCCTG	CAGGATGCCG	TGAGCAAGCT	750
GGACGACCCC	CTGCACCTGC	GCTGTACCAAG	CTCCCCAGAC	TACCTGGTGA	800
GCAGGGCCCCA	GGAGGCCTTG	GATGCCCTGA	GCACCTTGA	GGAGGGCCAC	850
GCCCAGTACC	TGACCTCCTT	GGCAGACGCC	TCCGCCCTCG	TGGCAGCTCT	900
GACCCGCTTC	TCCCACCTGG	CTGCCGATAC	CATCATCAAT	GGCGGTGCCA	950
CCTCGCACCT	GGCTCCCACC	GACCCTGCCG	ACCGCCTCAT	AGACACCTGC	1000
AGGGAGTGGG	GGGCCCCGGG	TCTGGAGCTC	ATGGGGCAGC	TGCAGGACCA	1050
GCAGGCTCTG	CGGCACATGC	AGGCCAGCCT	GGTGCGGACA	CCCCTGCAGG	1100
GCATCCTTCA	GCTGGGCCAA	GAAGTGAAC	CCAAGAGCCT	AGATGTGCGG	1150
CAGGAGGAGC	TGGGGGCCGT	GGTGACAAAG	GAGATGSCGG	CCACATCCGC	1200
AGCCATTGAA	GATGCTGTGC	GGAGGATTGA	GGACATGATG	AACCAGGCAC	1250
GCCACGCCAG	CTCGGGGCTG	AAGCTGGAGG	TGAACGAGAG	GATCCTCAAC	1300
TCTTGCACAG	ACCTGATGAA	GGCTATCCGG	CTCC'GGTGA	CGACATCCAC	1350
TAGCCTGCAG	AAGGAGATCG	TGGAGAGCGG	CAGGGGGGCA	GCCACGCAGC	1400
AGGAATTTTA	CGCCAAGAAC	TCGCGCTGGA	CCGAAGCCCT	CATCTCGGCC	1450
TCCAAGGCTG	TGGGCTGGGG	AGCCACACAG	CTGGTGGAGG	CAGCTGACAA	1500
GGTGTGCTT	CACACGGGCA	AGTATGAGGA	GCTCATCGTC	TGCTUCCACG	1550
AGATCGCAGC	CAGCACGGCC	CAGCTGGTGG	CGGCCTCCAA	GGTGAAGGCC	1600

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AACAAAGCACA GCCCCCACCTT GAGCCCCCTG CAGGAATGTT CTCGCACAGT 1650
CAATGAGAGG GCTGCCAATG TGGTGGCCTC CACCAAGTCA GGCCAGGAGC 1700
AGATTGAGCA CAGAGACACC ATGGATTTC TCGGCCTGTC CCTCATCAAG 1750
CTGAAGAGC AGGAGATGGA GACGCAGCTC CGTGTCTTGG AGCTGGAGAA 1800
GACGCTGGAG GCTGAACCCA TCGGCCTGGG GGAGTTGCGG AAGCAACACT 1850
ACGTGCTGGC TGGGGCATCA GGCAGCCCTG GAGAGGAGGT GGCCATCCGG 1900
CCCAGCACTG CCCCCCGAAG TGTAACCACC AAGAAACCAC CCCTGGCCCA 1950
GAAGCCCAGC GTGGCCCCCA GACAGGACCA CCAGCTTGAC AAAAAGGATG 2000
GCATCTACCC AGCTCAACTC GTGAACACT AGGCCCCCCA GGGGTCCAGC 2050
AGGGTGGCTG GTGACAGGCC TGGGCCTCTG CAACTGCCCT GACAGGACCG 2100
AGAGGCCCTG CCCCCTCCAC TGGTCCCCAA GCTTCCCGCC CCACCGTCTG 2150
GATCAATGTC CTCAGGCCCT CTGGCCCTTA CTGAGCCTGC AGGGTCCCTG 2200
GCCATGTGGG TGGTGTCTCT GGATGTGAGT CTCTTATTTA TCTGCAGAAG 2250
GAACCTTGGG GTGCAGCCAG GACCCGGTAG GCCTGAGCCT CAACTCTTCA 2300
GAAAATAGTG TTTTAAATAT TCCTCTTCAG AAAATAGTGT TTTTAAATAT 2350
CCGAGCTAGA GCTCTTCTTC CTACGTTTGT AGTCAGCACA CTGGGAAACC 2400
GGGCCAGCGT GGGCCTCCCT GCTTCTGGA CTCCTGAAGG TCGTGGATGG 2450
ATGGAAGGCA CACAGCCCGT GCTGGCTGAT GGGACGAGGG TCAGGCATCC 2500
TGCTGTGGC CTTCTGGGGC ACCGATTETA CCAGGCCCTC CAGCTGCGTG 2550
GTCTCCGCAG ACCAGGCTCT GTGTGGGCTA GAGGAATGTC GCCCATACC 2600
TCTTCAGGCC CTGGCCCTCG GGCCTCCGTG ATGGGAGCCC CCCAGGAGGG 2700
GTCAGATGCT GGAAGGGGCC GCTTCTTGGG GAGTGAGGTG AGACATAGCG 2750
GCCCAGBGC TGCCTTCACT CCTGGAGTTT CCATTTCAG CTGGAATCTG 2800
CAGCCACCCC CATTTCCCTGT TTTCCATTC CCCGTTCTGG CCGCGCCCCA 2850
CTGCCACCT CAAGGGGTGG TTTCCAGCCC TCCGGAGAGT GGGCTTGGCC 2900
CTAGGCCCTC CAGCTCAGCC AGAAAAAGCC CAGAAACCCA GGTGCTGGAC 2950
CAGGGCCCTC AGGGAGGGAC CCTGCGGCTA GAGTGGGCTA GGCCCTGGCT 3000
TTGCCCGTCA GATTGAACG AATGTGTGTC CCTTGAGCCC AAGGAGAGCG 3050
GCAGGAGGGG TGGGACCAGG CTGGGAGGAC AGAGCCAGCA GCTGCCATGC 3100
CCTCCTGCTC CCCCCACCCC AGCCCTAGCC CTTTAGCCCT TCACCCTGTG 3150
CTCTGGAAAG GCTACCAAAT ACTGGCCAAG GTCAGGAGGA GCAAAATGA 3200
GCCAGCACA GCGCTTGGC TTGTGTGTAG CATTTCCCTC TGAAGTGTTC 3250
TGTTGGCAAT AAAATGCACT TTCACTGTTA AAAAAAAAAA AAAAAAAAAA 3300
A

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(2) INFORMATION FOR SEQ ID NO: 7

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 676

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULAR TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(ix) FEATURE: Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:

Gly Glu Leu Glu Glu Gln Arg Lys Gln Lys Gln Lys Ala Leu Val

Asp	Asn	Glu	Gln	Leu	Arg	His	Glu	Leu	Ala	Gln	Leu	Arg	Ala	Ala	20	25	30
Gln	Leu	Glu	Arg	Glu	Arg	Ser	Gln	Gly	Leu	Arg	Glu	Glu	Ala	Glu	35	40	45
Arg	Lys	Ala	Ser	Ala	Thr	Glu	Ala	Arg	Tyr	Asn	Lys	Leu	Lys	Glu	50	55	60
Lys	His	Ser	Glu	Leu	Val	His	Val	His	Ala	Glu	Leu	Leu	Arg	Lys	65	70	75
Asn	Ala	Asp	Thr	Ala	Lys	Gln	Leu	Thr	Val	Thr	Gln	Gln	Ser	Gln	80	85	90
Glu	Glu	Val	Ala	Arg	Val	Lys	Glu	Gln	Leu	Ala	Phe	Gln	Val	Glu	95	100	105
Gln	Val	Lys	Arg	Glu	Ser	Glu	Leu	Lys	Leu	Glu	Glu	Lys	Ser	Asp	110	115	120
Gln	Gln	Glu	Lys	Leu	Lys	Arg	Glu	Leu	Glu	Ala	Lys	Ala	Gly	Glu	125	130	135
Leu	Ala	Arg	Ala	Gln	Glu	Ala	Leu	Ser	His	Thr	Glu	Gln	Ser	Lys	140	145	150
Ser	Glu	Leu	Ser	Ser	Arg	Leu	Asp	Thr	Leu	Ser	Ala	Glu	Lys	Asp	155	160	165
Ala	Leu	Ser	Gly	Ala	Val	Arg	Gln	Arg	Glu	Ala	Asp	Leu	Leu	Ala	170	175	180
Ala	Gln	Ser	Leu	Val	Arg	Glu	Thr	Glu	Ala	Ala	Leu	Ser	Arg	Glu	185	190	195
Gln	Gln	Arg	Ser	Ser	Gln	Glu	Gln	Gly	Glu	Leu	Gln	Gly	Arg	Leu	200	205	210
Ala	Glu	Arg	Glu	Ser	Gln	Glu	Gln	Gly	Leu	Arg	Gln	Arg	Leu	Leu	215	220	225
Asp	Glu	Gln	Phe	Ala	Val	Leu	Arg	Gly	Ala	Ala	Ala	Glu	Ala	Ala	230	235	240
Gly	Ile	Leu	Gln	Asp	Ala	Val	Ser	Lys	Leu	Asp	Asp	Pro	Leu	His	245	250	255
Leu	Arg	Cys	Thr	Ser	Ser	Pro	Asp	Tyr	Leu	Val	Ser	Arg	Ala	Gln	260	265	270

Glu	Ala	Leu	Asp	Ala	Val	Ser	Thr	Leu	Glu	Glu	Gly	His	Ala	Gln
				275					288					285
Tyr	Leu	Thr	Ser	Leu	Ala	Asp	Ala	Ser	Ala	Leu	Val	Ala	Ala	Leu
				290					295					300
Thr	Arg	Phe	Ser	His	Leu	Ala	Ala	Asp	Thr	Ile	Ile	Asn	Gly	Gly
				305					310					315
Ala	Thr	Ser	His	Leu	Ala	Pro	Thr	Asp	Pro	Ala	Asp	Arg	Leu	Ile
				320					325					330
Asp	Thr	Cys	Arg	Glu	Cys	Gly	Ala	Arg	Ala	Leu	Glu	Leu	Met	Gly
				335					340					345
Gln	Leu	Gln	Asp	Gln	Gln	Ala	Leu	Arg	His	Met	Gln	Ala	Ser	Leu
				350					355					360
Val	Arg	Thr	Pro	Leu	Gln	Gly	Ile	Leu	Gln	Leu	Gly	Gln	Glu	Leu
				365					370					375
Lys	Pro	Lys	Ser	Leu	Asp	Val	Arg	Gln	Glu	Glu	Leu	Gly	Ala	Val
				380					385					390
Val	Asp	Lys	Glu	Met	Ala	Ala	Thr	Ser	Ala	Ala	Ile	Glu	Asp	Ala
				395					400					405
Val	Arg	Arg	Ile	Glu	Asp	Met	Met	Asn	Gln	Ala	Arg	His	Ala	Ser
				410					415					420
Ser	Gly	Val	Lys	Leu	Glu	Val	Asn	Glu	Arg	Ile	Leu	Asn	Ser	Cys
				425					430					435
Thr	Asp	Leu	Met	Lys	Ala	Ile	Arg	Leu	Leu	Val	Thr	Thr	Ser	Thr
				440					445					450
Ser	Leu	Gln	Lys	Glu	Ile	Val	Glu	Ser	Gly	Arg	Gly	Ala	Ala	Thr
				455					460					465
Gln	Gln	Glu	Phe	Tyr	Ala	Lys	Asn	Ser	Arg	Trp	Thr	Glu	Gly	Leu
				470					475					480
Ile	Ser	Ala	Ser	Lys	Ala	Val	Gly	Trp	Gly	Ala	Thr	Gln	Leu	Val
				485					490					495
Glu	Ala	Ala	Asp	Lys	Val	Val	Leu	His	Thr	Gly	Lys	Tyr	Glu	Glu
				500					505					510
Leu	Ile	Val	Cys	Ser	His	Glu	Ile	Ala	Ala	Ser	Thr	Ala	Gln	Leu
				515					520					525

Val	Ala	Ala	Ser	Lys	Val	Lys	Ala	Asn	Lys	His	Ser	Pro	His	Leu
				530					535					540
Ser	Arg	Leu	Gln	Glu	Cys	Ser	Arg	Thr	Val	Asn	Glu	Arg	Ala	Ala
				545					550					555
Asn	Val	Val	Ala	Ser	Thr	Lys	Ser	Gly	Gln	Glu	Gln	Ile	Glu	Asp
				560					565					570
Arg	Asp	Thr	Met	Asp	Phe	Ser	Gly	Leu	Ser	Leu	Ile	Lys	Leu	Lys
				575					588					585
Lys	Gln	Glu	Met	Glu	Thr	Gln	Val	Arg	Val	Leu	Glu	Leu	Glu	Lys
				590					595					600
Thr	Leu	Glu	Ala	Glu	Arg	Met	Arg	Leu	Gly	Glu	Leu	Arg	Lys	Gln
				605					610					615
His	Tyr	Val	Leu	Ala	Gly	Ala	Ser	Gly	Ser	Pro	Gly	Glu	Glu	Val
				620					625					630
Ala	Ile	Arg	Pro	Ser	Thr	Ala	Pro	Arg	Ser	Val	Thr	Thr	Lys	Lys
				635					640					645
Pro	Pro	Leu	Ala	Gln	Lys	Pro	Ser	Val	Ala	Pro	Arg	Gln	Asp	His
				650					655					660
Gln	Leu	Asp	Lys	Lys	Asp	Gly	Ile	Tyr	Pro	Ala	Gln	Leu	Val	Asn
				665					670					675

Tyr

(2) INFORMATION FOR SEQ ID NO:8:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2338

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: mouse

(ix) FEATURE: cDNA for Huntingtin-interacting protein - mHIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:

GGCACGAGGG	CTCATTCAGA	TCCCCAGCT	GCCCGAGAAT	CCACCCAACTT	50
CCTACGAGCC	TCGGCCCTGT	CAGAGCACAT	CAGTCCTGTG	GTGGTGATCCC	100
GGCAGAGGTG	TCATCCCCAG	ACAGTGAGCC	TGTCCTGGAG	AAGGATGACCT	150
CATGGACATG	GACGCCTCCC	AGCAGACTTT	GTTTGACAAC	AACTTTCATGA	200

CGTCTTTGGC	AGCTCATTGA	GCAGCGACCC	TTTCAATTTT	AACAATCAAAA	250
TGGCGTGAAC	AAGGACGACA	AGGACCACTT	GATTGAACGC	CTGTACAGAGA	300
GATCAGTGGG	CTGACAGGUC	AGCTGGACAA	CATGAAGATT	GAGAGCCAGCG	350
GGCCATGCTG	CAGCTGAAGG	GTCGAGTGAG	TGAGCTGGAG	GCAGAGCTAGC	400
AGAGCAGCAG	CACTTGGGCC	GGCAGGCTAT	GGATGACTGC	GAGTTCTTGG	450
CACTGAGCTG	GATGAACTGA	AGAGGCAGCG	AGAGGACACG	GAGAAGGCACA	500
GGCAGCCTC	ACTGAGATAG	AAAGAAAGGC	CCAGGCTAAT	GAACAGAGGTA	550
TAGCAAGTTA	AAAGACAAGT	ACAGTGAACT	GGTCCAGAAC	CATCTCGACCT	600
GCTGCGGAAG	AACGCAGAGG	TGACCAAACA	GCTCTCCCTG	GCCCGCCAAAC	650
CCAGGTGGAT	TTGGAAAGAG	AGAAAAAGA	GCTAGCAGAT	TCCTTTGGCAC	700
GTGTAAGTGA	CCAGGCCACG	CGGAGACTC	AAGAGCAACA	GGATGTTCTA	750
GAGAACCCTG	AGCATGAACT	GGCCACCAGC	AGACAGGAGC	TGCAGGTCTT	800
CCACAGCAAC	CTGGAAACCT	CTGCCCAGTC	AGAAGCGAAA	TGGCTGACAC	850
AGATCCCCGA	GTTGGAGAAG	GAACAAGGCA	GCTTGGCGAC	TGTTGCAGCT	900
CAGAGAGAGC	AAGAGTTATC	AGCCCTCCGA	GACCAGCTGG	AAAGCACCCCA	950
GATCAAGCTG	GCTGCGGCC	AGGAATCCAT	GTGCCAGCAC	CTGAACGACC	1000
AGAGGAAAAC	CCTCTTGGCA	CCGATCAGGA	ACCTGCGGA	GCCTGAGATA	1050
CAGGAGGCGC	TGAGCCAGCT	TGAGGAACCC	ACCTTCATCA	GCTGTGCAGG	1100
ATCCACAGAT	CACCTTCTCT	CCAAAGTCAG	CTCCGTTCCT	AGCTGCCCTCG	1150
AGCAATGGA	AAAGAACGGC	AGCCAGTATC	TGGCCTGCC	AGAAGATATT	1200
AGTGAGCTTC	TGCACCTCGAT	CACCTTGCTT	GCCCCACTGA	CCGGTGACAC	1250
TGTCATCCAG	GGGAGTGCCA	CCAGCCTCCG	GGCCCCACCG	GAGCCAGCCG	1300
ACTCGTTGAC	CGAGCCCTGT	AGGCAGTATG	GCAGAGAAAC	CCTGGCCTAT	1350
CTGTCTCTCC	TGGAGGAAGA	GGGAACITGT	GAGAAITGCTG	ACGTCCACAGC	1400
CCTTAGGAAT	TGCCTCAGCA	CCCTCAAGAC	CCITGCGGAG	GAGCTGCTGC	1450
CCAGGGGCCCT	GGACATCAAG	CAGGAAGAGC	TGGGTGACCT	GGTGGACAAG	1500
GAGATGGCAG	CCACTTCAGC	TGCCATTGAA	GCTGCCACCA	CCCGGATAGA	1550
GGAAATTCTC	AGTAAGTCCC	GAGCAGGAGA	CACGGGAGTC	AAGCTGGAGG	1600
TGAATGAGAG	GATCCITGGGT	TCCTGTACCA	GCCTGATGCA	GGCCATCAAG	1650
GTGCTCGTTG	TGGCCTCCAA	GGACCTCCAG	AAGGAGATAG	TGGAGAGTGG	1700
CAGGGGTAGT	GCATCCCCCTA	AAGAAITTTA	CGCCAGAAAC	TCTCGGTGGA	1750
CGGAAGGGCT	GATATCCGCC	TCCAAACCTC	TTCGTITGGG	AGCTACCATC	1800
ATGGTGGATG	CTGCTGATCT	TGTGCTCCAA	GCCAAAGGCA	AGTTCGAGGA	1850
GCTGATCGTG	TGTTCAACGG	AGATTGCTGC	CAGTACTGCC	CAGCTCCTCC	1900
CTGCATCCAA	CGTGAAAGCG	AACAAGGGCA	GCCTCAATCT	GACCCAGCTG	2000
CAGCAGGCCT	CTCGAGGACT	GAACCAGGCC	ACAGCCGCTG	TGGTGGCCTC	2050
AACCATTTCT	GGCAATCTTC	AGATITGAGGA	AACAGACAGT	ATGGACTTCT	2100
CAAGCATGAC	ACTGACCCAG	ATCAAGCGCC	AGGAGATGGA	TTCCCAGGTIT	2150
AGGGTGCTGG	AGCTGGAAA	TGACCTGCAG	AAGGAGCGTC	AGAAACTAGG	2200
AGAGCTACGG	AAGAAACACT	ACGAGCTGGA	GGGCGTGGCT	GAGGGCTGGG	2250
AGGAAGGCAC	AGAAGCATCA	CCGTCTACTG	TCCAAGAAGC	AATACCGGAC	2300
AAAGAGTAGA	CCCAAGCCGA	CACCCACAC	ATCAGAAA		2338

(2) INFORMATION FOR SEQ ID NO: 9:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 676

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: mouse

(ix) FEATURE: Huntingtin-interacting protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9:

Ala	Arg	Gly	Leu	Ile	Gln	Ile	Pro	Gln	Leu	Pro	Glu	Asn	Pro	Pro	
				5					10					15	
Asn	Phe	Leu	Arg	Ala	Ser	Ala	Leu	Ser	Glu	His	Ile	Ser	Pro	Val	
				20					25					30	
Val	Val	Ile	Pro	Ala	Glu	Val	Ser	Ser	Pro	Asp	Ser	Glu	Pro	Val	
				35					40					45	
Leu	Glu	Lys	Asp	Asp	Leu	Met	Asp	Met	Asp	Ala	Ser	Gln	Gln	Thr	
				50					55					60	
Leu	Phe	Asp	Asn	Lys	Phe	Asp	Asp	Val	Phe	Gly	Ser	Ser	Leu	Ser	
				65					70					75	
Ser	Asp	Pro	Phe	Asn	Phe	Asn	Asn	Gln	Asn	Gly	Val	Asn	Lys	Asp	
				80					85					90	
Glu	Lys	Asp	His	Leu	Ile	Glu	Arg	Leu	Tyr	Arg	Glu	Ile	Ser	Gly	
				95					100					105	
Leu	Thr	Gly	Gln	Leu	Asp	Asn	Met	Lys	Ile	Glu	Ser	Gln	Arg	Ala	
				110					115					120	
Met	Leu	Gln	Leu	Lys	Gly	Arg	Val	Ser	Glu	Leu	Glu	Ala	Glu	Leu	
				125					130					135	
Ala	Glu	Gln	Gln	His	Leu	Gly	Arg	Gln	Ala	Met	Asp	Asp	Cys	Glu	
				140					145					150	
Phe	Leu	Arg	Thr	Glu	Leu	Asp	Glu	Leu	Lys	Arg	Gln	Arg	Glu	Asp	
				155					160					165	
Thr	Glu	Lys	Ala	Gln	Arg	Ser	Leu	Thr	Glu	Ile	Glu	Arg	Lys	Ala	
				170					175					180	
Gln	Ala	Asn	Glu	Gln	Arg	Tyr	Ser	Lys	Leu	Lys	Glu	Lys	Tyr	Ser	
				185					190					195	
Glu	Leu	Val	Gln	Asn	His	Ala	Asp	Leu	Leu	Arg	Lys	Asn	Ala	Glu	
				200					205					210	
Val	Thr	Lys	Gln	Val	Ser	Val	Ala	Arg	Gln	Ala	Gln	Val	Asp	Leu	
				215					220					225	
Glu	Arg	Glu	Lys	Lys	Glu	Leu	Ala	Asp	Ser	Phe	Ala	Arg	Val	Ser	

230					235					240				
Asp	Gln	Ala	Gln	Arg	Lys	Thr	Gln	Glu	Gln	Gln	Asp	Val	Leu	Glu
				245					250					255
Asn	Leu	Lys	His	Glu	Leu	Ala	Thr	Ser	Arg	Gln	Glu	Leu	Gln	Val
				260					265					270
Leu	His	Ser	Asn	Leu	Glu	Thr	Ser	Ala	Gln	Ser	Glu	Ala	Lys	Trp
				275					288					285
Leu	Thr	Gln	Ile	Ala	Glu	Leu	Glu	Lys	Gln	Gln	Gly	Ser	Leu	Ala
				290					295					300
Thr	Val	Ala	Ala	Gln	Arg	Glu	Glu	Glu	Leu	Ser	Ala	Leu	Arg	Asp
				305					310					315
Gln	Leu	Glu	Ser	Thr	Gln	Ile	Lys	Leu	Ala	Gly	Ala	Gln	Glu	Ser
				320					325					330
Met	Cys	Gln	Gln	Val	Lys	Asp	Gln	Arg	Lys	Thr	Leu	Leu	Ala	Gly
				335					340					345
Ile	Arg	Lys	Ala	Ala	Glu	Arg	Glu	Ile	Gln	Glu	Ala	Leu	Ser	Gln
				350					355					360
Leu	Glu	Glu	Pro	Thr	Leu	Ile	Ser	Cys	Ala	Gly	Ser	Thr	Asp	His
				365					370					375
Leu	Leu	Ser	Lys	Val	Ser	Ser	Val	Ser	Ser	Cys	Leu	Glu	Gln	Leu
				380					385					390
Glu	Lys	Asn	Gly	Ser	Gln	Tyr	Leu	Ala	Cys	Pro	Glu	Asp	Ile	Ser
				395					400					405
Glu	Leu	Leu	His	Ser	Ile	Thr	Leu	Leu	Ala	His	Leu	Thr	Gly	Asp
				410					415					420
Thr	Val	Ile	Gln	Gly	Ser	Ala	Thr	Ser	Leu	Arg	Ala	Pro	Pro	Glu
				425					430					435
Pro	Ala	Asp	Ser	Leu	Thr	Glu	Ala	Cys	Arg	Gln	Tyr	Gly	Arg	Glu
				440					445					450
Thr	Leu	Ala	Tyr	Leu	Ser	Ser	Leu	Glu	Glu	Glu	Gly	Thr	Val	Glu
				455					460					465
Asn	Ala	Asp	Val	Thr	Ala	Leu	Arg	Asn	Cys	Leu	Ser	Arg	Val	Lys
				470					475					480

Thr	Leu	Gly	Glu	Glu	Leu	Leu	Pro	Arg	Gly	Leu	Asp	Ile	Lys	Gln
				485					490					495
Glu	Glu	Leu	Gly	Asp	Leu	Val	Asp	Lys	Glu	Met	Ala	Ala	Thr	Ser
				500					505					510
Ala	Ala	Ile	Glu	Ala	Ala	Thr	Thr	Arg	Ile	Glu	Glu	Ile	Leu	Ser
				515					520					525
Lys	Ser	Arg	Ala	Gly	Asp	Thr	Gly	Val	Lys	Leu	Glu	Val	Asn	Glu
				530					535					540
Arg	Ile	Leu	Gly	Ser	Cys	Thr	Ser	Leu	Met	Gln	Ala	Ile	Lys	Val
				545					550					555
Leu	Val	Val	Ala	Ser	Lys	Asp	Leu	Gln	Lys	Glu	Ile	Val	Glu	Ser
				560					565					570
Gly	Arg	Gly	Ser	Ala	Ser	Pro	Lys	Glu	Phe	Tyr	Ala	Lys	Asn	Ser
				575					588					585
Arg	Trp	Thr	Glu	Gly	Leu	Ile	Ser	Ala	Ser	Lys	Ala	Val	Gly	Trp
				590					595					600
Gly	Ala	Thr	Ile	Met	Val	Asp	Ala	Ala	Asp	Leu	Val	Val	Gln	Gly
				605					610					615
Lys	Gly	Lys	Phe	Glu	Glu	Leu	Met	Val	Cys	Ser	Arg	Glu	Ile	Ala
				620					625					630
Ala	Ser	Thr	Ala	Gln	Leu	Val	Ala	Ala	Ser	Lys	Val	Lys	Ala	Asn
				635					640					645
Lys	Gly	Ser	Leu	Asn	Leu	Thr	Gln	Leu	Gln	Glu	Ala	Ser	Arg	Gly
				650					655					660
Val	Asn	Gln	Ala	Thr	Ala	Ala	Val	Val	Ala	Ser	Thr	Ile	Ser	Gly
				665					670					675
Lys	Ser	Gln	Ile	Glu	Glu	Thr	Asp	Ser	Met	Asp	Phe	Ser	Ser	Met
				680					685					690
Thr	Leu	Thr	Gln	Ile	Lys	Arg	Gln	Glu	Met	Asp	Ser	Gln	Val	Arg
				695					700					705
Val	Leu	Glu	Leu	Glu	Asn	Asp	Leu	Gln	Lys	Glu	Arg	Gln	Lys	Leu
				710					715					720
Gly	Glu	Leu	Arg	Lys	Lys	His	Tyr	Glu	Leu	Glu	Gly	Val	Ala	Glu
				725					730					735

Gly Trp Glu Glu Gly Thr Glu Ala Ser Pro Ser Thr Val Gln Glu
740 745 750

Ala Ile Pro Asp Lys Glu
755

(2) INFORMATION FOR SEQ ID NO:10:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 3964

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: cDNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: mouse

(ix) FEATURE: cDNA for Huntingtin-interacting protein - mHIP1a

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:

GGCAGGAGGC	GGCGCGCGGC	CTCCGTGTGC	CTAGGCTTGA	GGCGGGCGGT	50
GACGCCCTCAT	TCCGCGCGGAG	CCGGGCGCGG	ACACGCTCGC	CCCCAGCATC	100
AACAGCATCA	AGAAATGTGCC	GGCGCGGGTC	CTGAGCCGCA	GGCCGGGCCA	150
CAGCCTAGAG	GCCGAGCGCG	AGCACTTCCA	CAAGACGCAG	GCCATCAGTA	200
TCAGCAAAGC	CATCAACAGC	CAGGAGGCCC	CAGTGAAGGA	GAAGCATGCC	250
CGCGGTATCA	TCCTGGGCAC	GCATCATGAG	AAGGGAGCCT	TCACCTTCTG	300
GTCCATATGC	ATCGGCCTGC	CGCTGTCCAG	CAGCTCCATC	CTCAGCTGGA	350
AGTTCTGTCA	CGTCCTTCAC	AAGGTCCCTC	GGGACGGACA	CCCCAACGTC	400
CTGCATGACT	ATCAGCGGTA	CCGGAGCACC	ATACCTCACA	TCCGTGACTT	450
GTGGGGCCAC	CTTCGTGACC	AGTAATGGACA	CCTCGTCAAT	ATCTATACCA	500
AACTGTTGCT	GACTAAGATC	TCCCTCCACC	TTAAGCACCC	CCAGTTTCTT	550
GCAGGCCCTGG	AGGTAACAGA	TGACCTGTTG	GAGAAGGCGG	CGGGAACCTGA	600
TGTCAACAAC	ATTTTTCAGC	TTACCGTGGA	GATGTTTGAC	TACATGGACT	650
GTGAACTGAA	GCTTTCTGAG	TCAGTTTTC	GGCAGCTCAA	CACGGCCATC	700
GCAGTGTCCT	AGATGTCCTC	TGGCCAGTGT	CCCTAGCGC	CCCTCATCCA	750
GGTCATTCTG	GACTGCAGCC	ACCTGTATCA	CTACACAGTG	AAGCTCATGT	800
TTAAGCTGCA	CTCCTGTCTC	CCGGCAGACA	CCCTGCAAGG	CCACAGGGAT	850
CGGTTCCACC	ACCAGTTCCA	CAGCCTCAAA	AACCTCTTCC	GCCGGGCTTC	900
AGACATGCTG	TACTTCAAGA	GGCTCATCCA	GATCCCGCGG	CTGCCTGAGG	950
GACCCCCCAA	TTTCTTGGCG	GCTTCAGCCC	TGGCTGAGCA	CATCAAGCCG	1000
GTGGTGGTGA	TTCCCGAGGA	GGCCCCAGAG	GAAGAGGAGC	CTGAGAACCT	1050
AATTGAAATC	AGCAGTGCGC	CCCTGTCTGG	GGACCCACTC	GTCCCTGCTG	1100
ACCTCTTTGA	TCAGACCTTT	GGACCCCCCA	ATGGCTCCAT	GAAGGATGAC	1150
AGGGACCTCC	AAATCGAGAA	CTTGAAGAGA	GAGGTGGAGA	CCCTCCGTGC	1200
TGAGCTGGAG	AAGATTAAGA	TGGAGGCACA	CCCGTACATC	TCCAGCTGGA	1250
AGGGCCAGGT	GAATGGCCTG	GAGGCAGAGC	TGGAGGAGCA	GCCCAAGCAC	1300
AAGCAGAAGG	CCCTGGTGCA	CAACGAGCAG	CTGCGCCACG	AGCTGGCCCA	1350
GCTCAAGGCC	CTCCAGCTGG	AGGGCGCCCG	CAACCAGGGC	CTTCGAGAGG	1400
AAGCAGAGAG	GAAGGCCAGT	GCCACGGAGG	CACGCTACAG	CAAGCTGAAG	1450
GAGAAACACA	GCGAATCAT	TAACACGCAC	GCCGAGCTGC	TCAGGAAGAA	1500

CGCAGACACG	GCCAAGCAGC	TGACAGTGAC	ACAGCAGAGC	CAGGAGGAGG	1550
TGGCACGGCT	AAAGGAACAG	CTGGCCTTCC	AGATGGAGCA	AGCGAAGCGT	1600
GAGTCTGAGA	TGAAGATGGA	AGAGCAGAGC	GACCACTTGG	AGAAGCTCAA	1650
GAGGGAGCTG	GCGGCCAGGG	CACGAGAGCT	GGCCCCCTCC	CACGAGGGCC	1700
TGAGCCGCAC	AGAACACACT	GCCTCAGAGC	TCACTCACC	CCTCGACACA	1750
CTGAACGCGG	AGAAGGAAGC	CCTGACTGGA	CTCCTTCGGC	AGCGTGAGGC	1800
AGAGCTGCTG	GCGCTCAGA	GCCTCCTCCG	GGAGAAGGAG	GAGGCGCTTA	1850
GCCAAGAGCA	GCAGCGGAGC	TCCCAGGAGA	AGGGCGAGCT	ACGGGGGCAG	1900
CTGGCAGAAA	AGGAGTCTCA	GGAGCAGGGG	CTTCGGCAGA	AGCTGCTGGA	1950
TGAGCAGTIG	GCGGTGTTGC	GAAGTGCAGC	CGCCGAGGCA	GAGGCCATCC	2000
TACAGGATGC	AGTGAGCAAG	CTGGACGACC	CCCTGCACCT	CCGCTGCACC	2050
AGCTCCCCAG	ACTACTTGGT	GAGCCGGGCT	CAGGCAGCCC	TGGACAGCGT	2100
GAGCGGCTTG	GAGCAGGGCC	ACACCCAGTA	CCTGGCTTCC	TCCGAAGATG	2150
CTTCCTGCCCT	GGTGGCAGCG	CTGACCCGCT	TCTCCCATTT	GGCTGCCGAC	2200
ACCATTTGTC	ATGGTGGCGC	CACCTCCAC	CTGGCCCCCA	CCGACCCCGC	2250
CGACCGCCTG	ATGGACACAT	GCAGGGAGTG	TGGAGCCCGG	GCTCTGGAGC	2300
TGGTGGGACA	GCTGCAAGAC	CAGACAGTGC	TACGGAGGGC	TGAGCCGAGC	2350
CTGATGCCGG	CCCCCTGCA	GGGCATTCTG	CAGTTGGGCC	AGGACTTGAA	2400
GCCTAAGAGC	CTGGATGTAC	GGCAAGAGGA	GCTAGGGGCC	ATGGTGGACA	2450
AGGAGATGGC	GGCCACCTCG	GCAGCCATIG	AGGACGCTCT	CCCGAGGATC	2500
GAGGACATGA	TGAGCCAGGC	CCGCCACGAC	AGCTCAGGCG	TGAAACTGCA	2550
GGTGAATGAG	AGGATCCTCA	ACTCCTGCAC	AGACCTGATG	AAGGCTATCC	2600
GGCTCCTGGT	GATGACCTCC	ACCAGCCTGC	AGAAGGAAT	TGTGGAGAGC	2650
GGCAGGGGGC	CAGCAACGCA	GCAGGAATTT	TATGCCAAGA	ATTACCGGTG	2700
GACTGAAGGC	CTCATCTCAG	CCTCTAAGGC	AGTGGGCTGG	GGAGCCACAC	2750
AGCTGGTGGG	GTCACCTGAC	AAGGTGTGTC	TTCACATGGG	CAABTACGAG	2800
GAACTCATCG	TCTCCTCCCA	TGAGATTGCG	GCCAGCACGG	CCAGCTGGT	2850
GGCAGCCTCG	AAGGTGAAG	CCAACAAGAA	CAGTCCCCAC	TTGAGCCGCG	2900
TGCAGGAATG	TTCCCGCACT	GTCAACGAGA	GGGCTGCCAA	CGTCGTGGCC	2950
TCCACCAAAI	CTGGCCAGGA	GCAGATTGAG	GACAGAGACA	CCATGGATTT	3000
CTCTGGCCTC	TCCCTCATCA	AGTTGAAGAA	GCAGGAGATG	GAGACACAGC	3050
TGCGAGTCTT	GGAGCTGGAG	AAGACACTAG	AGGCAGAGCG	TGTCCGGCTC	3100
GGGGAGCTTC	GGAAACAGCA	CTATGTACTG	GCTGGGGGGA	TGGGAACACC	3150
TAGCGAAGAA	GAACCCAGCA	GACCCAGCCC	AGCTCCCCGA	AGTGGGGCCA	3200
CTAAGAAGCC	ACCGCTGGCC	CAGAAACCCA	CCATACCCCC	CAGGACAGAC	3250
AACCAGCTCGA	CAAAAAGGAT	GGTGTCTACU	CAGCTCAACT	TCTGAACCTAC	3300
TAGGCCCCCTAA	GGTGTTCAGC	AGGATGGCTG	GTGGTTGTGC	CTGCCCTTCA	3350
TGTGCTGTCT	GGCAGTGGTC	AAGGGGCTTC	TGAGAAGCCT	CCAACCTCTC	3400
CCCAAGGGGCC	TAGTCTGTGG	GACAGTTCAT	CTGGATGTGA	ATCTATTTAT	3450
CTTAAGTAGGA	ACTGCCCTCGA	CCAGCTGGGA	CCCAGCAGGC	CTGAGCCACA	3500
AACTCTGACCG	GACATCAGAG	ATAGTCTGAA	TCTTCCGAGG	TATTTCTTTC	3550
TTCGTAAGTTI	AGTCAGCACA	CTGGGAAGAG	GTCACATAAG	CCAGGAGCCT	3600
CCTTGTCTCTG	GACTCAAAAG	TCTGAGGCCCT	TAAGTGAACA	ACAGAAAGAG	3650
GGTCCCTGCTG	GCTACCAAGG	ATAAGGGGAT	GACCTGTGAC	CCTTGAGCCA	3700
GGGAGAGCAGG	TAACTGGCTT	GGTGTCTATCA	CCTGGGGGCC	TGGTGTCTAGG	3750
GCATCCATGCT	GGGAGCCCCA	GCAGACCAGG	CTTTGTGTTC	GAGCCTGGCA	3800
TCATCGTGGCT	GGGGCAGCCC	CTGCTCAGCT	CCTGTCTCTC	CCCGTGACCT	3850
TGAAGCCACCC	TCCCCCGGTA	CAGTTTTCCA	TTCTCTTGGC	TACTAGTGTG	3900
GCTGTTCAATG	CCTACCTTGA	TGAGTAGATT	TCAGCCCTCC	TAAAGCTGGG	3950
GCCTTTCTCTG	TGCC				3964

(2) INFORMATION FOR SEQ ID NO: 11:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 676

(B) TYPE: protein

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein

(iii) HYPOTHETICAL: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: mouse

(ix) FEATURE: Huntingtin-interacting protein -mHIP1a

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:

Met	Asn	Ser	Ile	Lys	Asn	Val	Pro	Ala	Arg	Val	Leu	Ser	Arg	Arg	5	10	15
Pro	Gly	His	Ser	Leu	Glu	Ala	Glu	Arg	Glu	Gln	Phe	Asp	Lys	Thr	20	25	30
Gln	Ala	Ile	Ser	Ile	Ser	Lys	Ala	Ile	Asn	Ser	Gln	Glu	Ala	Pro	35	40	45
Val	Lys	Glu	Lys	His	Ala	Arg	Arg	Ile	Ile	Leu	Gly	Thr	His	His	50	55	60
Glu	Lys	Gly	Ala	Phe	Thr	Phe	Trp	Ser	Tyr	Ala	Ile	Gly	Leu	Pro	65	70	75
Leu	Ser	Ser	Ser	Ser	Ile	Leu	Ser	Trp	Lys	Phe	Cys	His	Val	Leu	80	85	90
His	Lys	Val	Leu	Arg	Asp	Gly	His	Pro	Asn	Val	Leu	His	Asp	Tyr	95	100	105
Gln	Arg	Tyr	Arg	Ser	Asn	Ile	Arg	Glu	Ile	Gly	Asp	Leu	Trp	Gly	110	115	120
His	Leu	Arg	Asp	Gln	Tyr	Gly	His	Leu	Val	Asn	Ile	Tyr	Thr	Lys	125	130	135
Leu	Leu	Leu	Thr	Lys	Ile	Ser	Phe	His	Leu	Lys	His	Pro	Gln	Phe	140	145	150
Pro	Ala	Gly	Leu	Glu	Val	Thr	Asp	Glu	Val	Leu	Glu	Lys	Ala	Ala	155	160	165
Gly	Thr	Asp	Val	Asn	Asn	Ile	Phe	Gln	Leu	Thr	Val	Glu	Met	Phe	170	175	180
Asp	Tyr	Met	Asp	Cys	Glu	Leu	Lys	Leu	Ser	Glu	Ser	Val	Phe	Arg			

					185					190					195
Gln	Leu	Asn	Thr	Ala	Ile	Ala	Val	Ser	Gln	Met	Ser	Ser	Gly	Gln	
				200					205					210	
Cys	Arg	Leu	Ala	Pro	Leu	Ile	Gln	Val	Ile	Gln	Asp	Cys	Ser	His	
				215					220					225	
Leu	Tyr	His	Tyr	Thr	Val	Iys	Leu	Met	Phe	Lys	Leu	His	Ser	Cys	
				230					235					240	
Leu	Pro	Ala	Asp	Thr	Leu	Gln	Gly	His	Arg	Asp	Arg	Phe	His	Glu	
				245					250					255	
Gln	Phe	His	Ser	Leu	Iys	Asn	Phe	Phe	Arg	Arg	Ala	Ser	Asp	Met	
				260					265					270	
Leu	Tyr	Phe	Lys	Arg	Leu	Ile	Gln	Ile	Pro	Arg	Leu	Pro	Glu	Gly	
				275					288					285	
Pro	Pro	Asn	Phe	Leu	Arg	Ala	Ser	Ala	Leu	Ala	Glu	His	Ile	Lys	
				290					295					300	
Pro	Val	Val	Val	Ile	Pro	Glu	Glu	Ala	Pro	Glu	Glu	Glu	Glu	Pro	
				305					310					315	
Glu	Asn	Leu	Ile	Glu	Ile	Ser	Ser	Ala	Pro	Pro	Ala	Gly	Glu	Pro	
				320					325					330	
Val	Val	Val	Ala	Asp	Leu	Phe	Asp	Gln	Thr	Phe	Gly	Pro	Pro	Asn	
				335					340					345	
Gly	Ser	Met	Lys	Asp	Asp	Arg	Asp	Leu	Gln	Ile	Glu	Asn	Leu	Lys	
				350					355					360	
Arg	Glu	Val	Glu	Thr	Leu	Arg	Ala	Glu	Leu	Glu	Lys	Ile	Lys	Met	
				365					370					375	
Glu	Ala	Gln	Arg	Tyr	Ile	Ser	Gln	Leu	Lys	Gly	Gln	Val	Asn	Gly	
				380					385					390	
Leu	Glu	Ala	Glu	Leu	Glu	Glu	Gln	Arg	Lys	Gln	Lys	Gln	Lys	Ala	
				395					400					405	
Leu	Val	Asp	Asn	Glu	Gln	Leu	Arg	His	Glu	Leu	Ala	Gln	Leu	Lys	
				410					415					420	
Ala	Leu	Gln	Leu	Glu	Gly	Ala	Arg	Asn	Gln	Gly	Leu	Arg	Glu	Glu	
				425					430					435	
Ala	Glu	Arg	Lys	Ala	Ser	Ala	Thr	Glu	Ala	Arg	Tyr	Ser	Lys	Leu	

				440					445					450
Lys	Glu	Lys	His	Ser	Glu	Leu	Ile	Asn	Thr	His	Ala	Glu	Leu	Leu
				455					460					465
Arg	Lys	Asn	Ala	Asp	Thr	Ala	Lys	Gln	Leu	Thr	Val	Thr	Gln	Gln
				470					475					480
Ser	Gln	Glu	Glu	Val	Ala	Arg	Val	Lys	Gln	Gln	Leu	Ala	Phe	Gln
				485					490					495
Met	Gln	Gln	Ala	Lys	Arg	Glu	Ser	Glu	Met	Lys	Met	Glu	Glu	Gln
				500					505					510
Ser	Asp	Gln	Leu	Glu	Lys	Leu	Lys	Arg	Glu	Leu	Ala	Ala	Arg	Ala
				515					520					525
Gly	Glu	Leu	Ala	Arg	Ala	Gln	Glu	Ala	Leu	Ser	Arg	Thr	Glu	Gln
				530					535					540
Ser	Gly	Ser	Glu	Leu	Ser	Ser	Arg	Leu	Asp	Thr	Leu	Asn	Ala	Glu
				545					550					555
Lys	Gln	Ala	Leu	Ser	Gly	Val	Val	Arg	Gln	Arg	Glu	Ala	Glu	Leu
				560					565					570
Leu	Ala	Ala	Gln	Ser	Leu	Val	Arg	Glu	Lys	Glu	Glu	Ala	Leu	Ser
				575					580					585
Gln	Glu	Gln	Gln	Arg	Ser	Ser	Gln	Glu	Lys	Gly	Glu	Leu	Arg	Gly
				590					595					600
Gln	Leu	Ala	Glu	Lys	Glu	Ser	Gln	Glu	Gln	Gly	Leu	Arg	Gln	Lys
				605					610					615
Leu	Leu	Asp	Glu	Gln	Leu	Ala	Val	Leu	Arg	Ser	Ala	Ala	Ala	Glu
				620					625					630
Ala	Glu	Ala	Ile	Leu	Gln	Asp	Ala	Val	Ser	Lys	Leu	Asp	Asp	Pro
				635					640					645
Leu	His	Leu	Arg	Cys	Thr	Ser	Ser	Pro	Asp	Tyr	Leu	Val	Ser	Arg
				650					655					660
Ala	Gln	Ala	Ala	Leu	Asp	Ser	Val	Ser	Gly	Leu	Glu	Gln	Gly	His
				665					670					675
Thr	Gln	Tyr	Leu	Ala	Ser	Ser	Glu	Asp	Ala	Ser	Ala	Leu	Val	Ala
				680					685					690
Ala	Leu	Thr	Arg	Phe	Ser	His	Leu	Ala	Ala	Asp	Thr	Ile	Val	Asn

				695					700					705
Gly	Ala	Ala	Thr	Ser	His	Leu	Ala	Pro	Thr	Asp	Pro	Ala	Asp	Arg
				710					715					720
Leu	Met	Asp	Thr	Cys	Arg	Glu	Cys	Gly	Ala	Arg	Ala	Leu	Glu	Leu
				725					730					735
Val	Gly	Gln	Leu	Gln	Asp	Gln	Thr	Val	Leu	Arg	Arg	Ala	Gln	Pro
				740					745					750
Ser	Leu	Met	Arg	Ala	Pro	Leu	Gln	Gly	Ile	Leu	Gln	Leu	Gly	Gln
				755					760					765
Asp	Leu	Lys	Pro	Lys	Ser	Leu	Asp	Val	Arg	Gln	Glu	Glu	Leu	Gly
				770					775					780
Ala	Met	Val	Asp	Lys	Glu	Met	Ala	Ala	Thr	Ser	Ala	Ala	Ile	Glu
				785					790					795
Asp	Ala	Val	Arg	Arg	Ile	Glu	Asp	Met	Met	Ser	Gln	Ala	Arg	His
				800					805					810
Glu	Ser	Ser	Gly	Val	Lys	Leu	Glu	Val	Asn	Glu	Arg	Ile	Leu	Asn
				815					820					825
Ser	Cys	Thr	Asp	Leu	Met	Lys	Ala	Ile	Arg	Leu	Leu	Val	Met	Thr
				830					835					840
Ser	Thr	Ser	Leu	Gln	Lys	Glu	Ile	Val	Glu	Ser	Gly	Arg	Gly	Ala
				845					850					855
Ala	Thr	Gln	Gln	Glu	Phe	Tyr	Ala	Lys	Asn	Ser	Arg	Trp	Thr	Glu
				860					865					870
Gly	Leu	Ile	Ser	Ala	Ser	Lys	Ala	Val	Gly	Trp	Gly	Ala	Thr	Gln
				875					880					885
Leu	Val	Glu	Ser	Ala	Asp	Lys	Val	Val	Leu	His	Met	Gly	Lys	Tyr
				890					895					900
Glu	Glu	Leu	Ile	Val	Cys	Ser	His	Glu	Ile	Ala	Ala	Ser	Thr	Ala
				905					910					915
Gln	Leu	Val	Ala	Ala	Ser	Lys	Val	Lys	Ala	Asn	Lys	Asn	Ser	Pro
				920					925					930
His	Leu	Ser	Arg	Leu	Gln	Glu	Cys	Ser	Arg	Thr	Val	Asn	Glu	Arg
				935					940					945
Ala	Ala	Asn	Val	Val	Ala	Ser	Thr	Lys	Ser	Gly	Gln	Glu	Gln	Ile

	950		955		960
Glu Asp Arg Asp Thr Met Asp Phe Ser Gly Leu Ser Leu Ile Lys					
	965		970		975
Leu Lys Lys Gln Gln Met Glu Thr Gln Val Arg Val Leu Glu Leu					
	980		985		990
Glu Lys Thr Leu Glu Ala Glu Arg Val Arg Leu Gly Glu Leu Arg					
	995		1100		1105
Lys Gln His Tyr Val Leu Ala Gly Gly Met Gly Thr Pro Ser Glu					
	1110		1115		1120
Glu Glu Pro Ser Arg Pro Ser Pro Ala Pro Arg Ser Gly Ala Thr					
	1125		1130		1135
Lys Lys Pro Pro Leu Ala Glu Lys Pro Ser Ile Ala Pro Arg Thr					
	1140		1145		1150
Asp Asn Gln Leu Asp Lys Lys Asp Gly Val Tyr Pro Ala Gln Leu					
	1155		1160		1165
Val Asn Tyr					

(2) INFORMATION FOR SEQ ID NO:12:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 18

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: other DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12:

GAAGATACCC CACCAAAC 18

(2) INFORMATION FOR SEQ ID NO:13:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 35

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: other DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13:

GCTTGACAGT GTAGTCATAA AGGTGGCTGC AGTCC 35

(2) INFORMATION FOR SEQ ID NO:14:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 24

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: other DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14:

GGACATCTCC AGGGAGTTGA ATAC 24

(2) INFORMATION FOR SEQ ID NO:15:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 41

(B) TYPE: nucleic acid

(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: other nucleic acid

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: yes

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15:

CUACUACUAC UACUAGGCCA CGGTCGACT AGTACGGGII GGGGGGGII G 41

(2) INFORMATION FOR SEQ ID NO:16:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 516

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 1 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:

TCTGTGGAG	GTCTGGAGG	SAGAGAGGG	CAGCTGGATG	CTCTTGGGC	AUGGTGGGC	60
CTGATCTCT	CGCTCTTCC	TCTGTCTCT	GGAGAAATA	TGTTTCCCT	GGGGATGAA	120
GCATCTCTT	GTGCGGGCT	TAAATGCCAT	GTGTGTGTG	CAAGGGAGT	AGTGGCGAG	180
GGACCAAC	CTGGCACAC	CCAAATGCC	GCAGTCTTT	CCACTCCCT	ACCAAGGCA	240
GGCACTG	TCCTGGGAG	GCTGCCCC	TCTGCCCTC	GCTGGGCGC	TGCAAGGAC	300
CGACCACT	TGGGGCTGG	GAACCTTGG	TGGAGCAG	GAAAGGCGG	GGCTCTGGA	360
GGGTCTC	CACCTCTCA	GGCTTATTC	TCTCATCTC	CTTTCCCTC	CCCTTCTGT	420
TTTTCTG	ACTAGCATC	ATAAGGCAT	TAAATGCG	GAAGTGGCT	TAAAGGAAA	480
ACACGCCA	AATATCTTT	GGATGTTGT	TGGAG			516

(2) INFORMATION FOR SEQ ID NO:17:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 193

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(v) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 2 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17:

TGTTTTCCAT	AACCCCCCT	CACCGTCAT	ACTGGGCACC	CACCATGAGA	AAGGGGCACA	60
GACCTTCTG	TCTGTTGTCA	ACCGCTGCC	TCTGTCTAGC	AACTCAATGC	TCTCTCTGCA	120
GTCTGTCCAT	GTCTTCCACA	AACCTCTCC	AGATGGACAC	CCGACCTGA	GTCTCTGGG	180
CTATGGCTG	GCA					193

(2) INFORMATION FOR SEQ ID NO:18:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 104

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(v) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 3 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:

GTGTTCTTT	GCCCCGAC	CTCTGAGG	ACTCTCTGAG	ATACAGAAAT	GAATTGAGTG	60
ACATGAGCAG	GATGTGGTG	AGTTTGGAGA	TGTAATCAGG	AGCC		104

(2) INFORMATION FOR SEQ ID NO:20:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 327
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: double
 (D) TOPOLOGY: linear
 (ii) MOLECULE TYPE: genomic DNA
 (iii) HYPOTHETICAL: no
 (iv) ANTI-SENSE: no
 (vi) ORIGINAL SOURCE:

(A) ORGANISM: human
 (x) FEATURE: exon 4 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:

AATTCCTGGC	TCCAGATCTC	TTGACTGTTA	TGTTCTTCTT	CTTCACTCTC	TTTCCCTTTC	60
TCTTCTTAAA	AGGGCCACCT	GAGCGAGGGG	TATGGCCAGC	TGTGCAGCCT	CTACCTGAAA	120
CTGCTAAGAA	CCAAGATCCA	GTACCACACC	AAAGTGACTC	TCTTCCGACA	CTTCTGCCGC	180
CACCCGCGCC	TCCCTGCTC	CATCCCTTCA	GGCCCTCCCT	GGGCTCATT	GTGAGCTCTT	240
TCAGGTAATA	GACAGCCCA	CTTCTGAGG	AAGTCTCCAC	ATCATCTACC	CAAGCTGTCA	300
GAGAGCAANG	CCACCGCCAG	GCCCAAG				327

(2) INFORMATION FOR SEQ ID NO:21:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 331
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: double
 (D) TOPOLOGY: linear
 (ii) MOLECULE TYPE: genomic DNA
 (iii) HYPOTHETICAL: no
 (iv) ANTI-SENSE: no
 (vi) ORIGINAL SOURCE:

(A) ORGANISM: human
 (x) FEATURE: exon 5 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:

GGGCTCAAGC	AATCTTCCCA	CCTCGGCTC	CCAGTAGCT	GGGACCACAG	GCTGTGCCA	60
CCACGCCCCG	CTGAGAGAGG	GGCTTCATG	TCTTCTGCCC	TGACTCCCTT	CTTCTGCTC	120
CCCTCCAGAA	TCCCACTTC	CCAGSCACC	TGCAGATGAG	TGACCGCCAG	CTGGACGAGG	180
CTCGAGAAAG	TGACCTGAAC	AACCTGTAA	TGGCTCTTGC	CCGAGGCCA	GGGAGGAGCA	240
AAGCTTTTGT	GAATGCTGAC	ACTTCTCATA	AGCTTCTTCC	AGCCCTCAT	GGGGGGAGGC	300
CGTGGCTGGG	ATGCGCAGCA	AGGCTCTGG	G			331

(2) INFORMATION FOR SEQ ID NO:22:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 470
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: double
 (D) TOPOLOGY: linear
 (ii) MOLECULE TYPE: genomic DNA
 (iii) HYPOTHETICAL: no
 (iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 6 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 22:

ACTGTCGCTG	TCACTGTTGA	CTTCACCAGG	CTGATGBCU	ATAATACCCA	CAAGGCTAAG	60
ACTTGGAGCT	GGAGTTCTGT	GTGTCTTTTC	CCATCCACAT	GAGCATTGGA	GACTGGAGTA	120
CCCTAGAGCG	TGGGGGAGGG	GACAGGTAAC	AGACCGCCCT	CAGSCTGTGG	AGTGTAAAGCT	180
CTCTTTTCTC	TJGGGTTCAG	TTTCCAGTAA	ACAGTTCAGA	TGTTTGACTA	CCCTGGAGTCT	240
CAACTCAACC	TCTTCCAAAC	AGGTGAGTCT	CTTCCCTCCC	GTCTAACCCA	GGCTCTCATG	300
GGACTACCT	AATTCCTAGT	CTCTCTCTCC	CTGCAAGATG	TGCAACACAA	GGGGTAGGAA	360
AATCCAGACA	TTACACACCC	ATCTCTGGTC	TCTCCAACCC	TCTGTGAGGG	AGGGACTGAA	420
CTCTTCTAGT	ATTTTCTCTT	TTAAGAGACA	AGTCTCTGGC	CGGGTGCAGT		470

(2) INFORMATION FOR SEQ ID NO:23:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 565

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 7 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23:

TCTTCACCTG	TTTAAAGGGG	ATACCTTTAC	CTATCTCATG	GGAGTGTGTG	GAAGGTTAA	60
TCAATTAGAT	GAGGTAAAGC	ACGCACAGAA	TGGTCTCTTG	GTGTATGTTC	GAUCCCTGCC	120
TCCTGCCCCC	TGAAGAGCC	CCCTCTAATC	CCCTGGCTCT	ACCACTTTTC	TCCCTCACTT	180
TTATTCTCTA	GTATTCAACT	CCCTGGACAT	GTCTTCTCTT	GTCTTCTCTA	CGGCAACAGC	240
GCAGTGGGCG	CTGCCCCCGC	TGATCCAGCT	CATCTTTGAC	TCTAGCCACC	TTATGACTTA	300
CACTCTCAGC	CTTCTCTCTA	AATCCACTC	CTGTGAGTAC	CGCGGGCCAG	ATCTTCTTAC	360
ATGAGATTCA	GGCCAGAGGG	AGGATCCAG	CTTGAGGATG	TCTCCAGAGA	AACCCACTTC	420
TTCTCAGTGC	CTTTGGCTCT	CTGCTCTCTT	TCCAAANGGC	CCCGGAGCTT	CTGACCATTG	480
TGACCTTAAA	AGAGCAGGGC	CCAGGCTTTG	GTGACCCCGC	TAAAGCCCCC	GGCTTGGCCG	540
TCTTGGCTCC	AGTGTTCAGC	GATCT				565

(2) INFORMATION FOR SEQ ID NO:24:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 233

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 8 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 24:

GGGACAGCTC	TAGGCCAGTC	CTTCCGCTG	GCAGTCTG	CCACATGCCC	CAGGGTAGCT	60
GGGCCCCCTC	CCCTCGAGAG	CCCCGCTGTG	GCTTCCCTGC	CCTCTGGTCC	CCCTCCCCCTC	120
TCACACCTCT	TCCAATTTCT	TCCAGCCCTC	CCAGCTGACA	CCCTGCAAGG	CCACCGGGAG	180
CGCTTCATGG	AGCAGTTTAC	AAAGTAAGTG	GTTCAAGTAA	CAGGAATGGA	GGT	233

(2) INFORMATION FOR SEQ ID NO: 25:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 578

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exons 9 and 10 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 25:

TGAATCCAG	CACCATGGAG	TTTATCTCT	TGACAGCCCTG	TCCCTTTGGG	CTGGGGAGCC	60
GGCAGGAAAG	CCAGGTGGCT	GCTCTGTCCC	CTACATGGGG	CTGATGAAGA	CACCCAGCAC	120
CCCTCAGGTC	TTTCTTCCAC	CCTAGGTGA	AGGATCTGTT	CTACCGCTTC	AGCAGCTCC	180
AGTACTTCAA	GCGGCTCAT	CAGATCCCC	AGCTGCCTGA	GGTAAGCATG	CCCAACCACA	240
CACCCCTCGC	ATTGACAGAG	CCCAAGGTAC	TCTCTTAAGG	GGGGGGGGGG	CTTGGCAAGC	300
AGGCACTATT	TGAGGATGAG	TCTCCGTCTT	CAGAACCCAC	CCAACCTTCT	GCGAGCTTCA	360
GGCTGTCTAG	AACATATCAG	CCCTGTGGTG	GTGATCCCTG	CAGAGGCCCTC	ATCCCCCGAC	420
AGCGAGGAC	TCTAGAGCAA	CCATGACCTC	ATGGACATCG	ATGCTCTCTA	GCAGGTGAGG	480
ACCACTTCCC	AGAGAACTT	GGCTTTCTCT	CTCAGCTGCA	AGTACAGGGG	AGAGGCTGGG	540
GGGACCCCTG	GCCAAAGCCC	ATTGACTCTA	ACCAGCTT			578

(2) INFORMATION FOR SEQ ID NO: 26:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 390

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 11 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 26:

AAAAAATTT	AAAAAAATTA	ACAAGTCTCA	ACCTTTAAT	TCCAGAAAGC	GGGCATTCTC	60
CCATATCACT	CAACTGACCC	ACACACAGAA	TTCTCTGGCT	CTCTGACTTA	TTCTCACTCC	120
TTTTTGGTCA	ACCACAGAT	TTATTGACA	ACAAGTTTGA	TGACATCTTT	CCCACTTCAT	180
TCAGCAGTGA	TCCCTTCAAT	TTCACAGCTC	AAAATCTCTT	CAACAGGAT	GAGAGGTGAG	240
TCCAAGCTGG	GTTCAAGCAC	ATCTTTAGG	AGCTAAGTTA	AGCCATGGTC	TGCTTCAAAA	300
CACTAACCAA	AGAGGATTC	TTAATGATA	TGGGGCTTCT	TAGATACAGA	ACATCTTCAA	360
GGGTTGGGGG	CAATGCTTA	TGCTTCAAT				390

(2) INFORMATION FOR SEQ ID NO:27:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 547

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 12 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:

AAATCAATA	ACCATGGATT	TATGAGTACT	AGATTAGTAT	CTGGTAACAT	TTAGAGTATA	50
ATTATGSCA	TTTCAAGCAA	TTGTCCCCAA	ATTANTACCA	GCTTTTAATT	TCCTCCTCTG	120
AGCTCACAAT	TAAAAACAGA	GGGATAGAAG	CACATAGAAA	GCAAACTCAT	TCCCCCTCTC	180
TTCCAGUGA	CCACTTAATT	GAGCGACTAT	ACAGAGAGAT	CACTCGATTG	AAGCCACAGC	240
TAGAAACAT	GAAGACTGAG	GTATAACTTG	GATCTGCTCT	GCCTTTGGGC	TCACCCAAA	300
CAGGATAGAT	TTGAATCTTA	AATTGCAATC	ACACTAGCCA	CCCACACTTG	CTCACACCTG	360
TAATCTTAGC	ACTTTGGGAG	GCCAAAGGCG	GAGGATTACC	TGAGGTGGGG	AGTTCGAGAC	420
CAGCTTGGGC	AACAGGCTGA	AACCCCCCTC	TTCAATAAAA	ATGCATTAAT	TAGCCGGGTG	480
TSTTGGCAGG	CACCTCTAAT	CCAGCTACT	CGGGAAGCTG	AGGCATGAGA	ATTGCTTGAA	540
CTTGGGA						547

(2) INFORMATION FOR SEQ ID NO:28:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 436

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 13 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 28:

CCCCCAGCCA	CTCTAAGAG	GACCACAAAT	CCCCGGGCAT	CATCCCCCTGT	TATTGTTGTT	50
GATTGAGGGG	CTCTAATGA	CCAGATGGTC	CAACCTCTCT	CCCACCTGCA	GAGTTGACTT	120
AGGGGAATCA	GCTATTTACT	TGGAGCATAG	GTAGGACCCG	CTTCTCCGGC	CCATGCCCTGT	180
GACCCGTGGC	AGTGGGCGGT	TGGCCTCATG	ACCGGAGTCT	CCCCACAGAG	CCAGXXGGGT	240
GTGCTGCAGC	TGAAGGUGCA	CGTCAGCGAG	CTGGAAGCAC	ATCTCCCCCA	GCATCAGCAC	300
CTGCGGCAGC	AGCCCCCCCA	CAACTGTGAA	TTCTTCTGGG	CAGAACTGGA	CGAGCTCAGG	360
AGGCAGTGGG	AGGACACCGA	GAGGCTTACG	CGGAGCTCTT	CTGAATAGCA	AAGTCACCCC	420
TGGGTGGGGG	CGGGGG					436

(2) INFORMATION FOR SEQ ID NO:29:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 469
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: double
 (D) TOPOLOGY: linear
 (ii) MOLECULE TYPE: genomic DNA
 (iii) HYPOTHETICAL: no
 (iv) ANTI-SENSE: no
 (vi) ORIGINAL SOURCE:
 (A) ORGANISM: human
 (x) FEATURE: exon 14 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:

GACTTGAGGC	CAAGGAGTC	AAGGCTGCAG	TGAACACTGA	TTCTCCCACT	CCACCCACAG	60
CTGCCTGACA	GAGCAAGACT	GTCTCATAAC	AAAACAAGGA	GGACCTTCTA	GGGACCTTGG	120
CTCATTTCAA	GGAAGGCAAG	GCTCCCTGCT	AGCTTAGACT	CTTCACCTTC	CTCCTTTACA	180
ATACAGGGA	AGCTCAAGCC	ATGAACAGC	GATATAGCAA	GCTAAAGGAG	AAGTACAGCG	240
AGCTGCTTCA	GAACACGCT	GACCTGCTGC	GGAAGGTAG	ACCTCAGCC	CCTGTCACCA	300
TCTTGCAGGC	CTTCACCTTC	TAGGGAGACA	GCGGCTCAGG	CTGTGCTT	CCCCGGGGCC	360
AGCAACCCCT	ACATTGATCT	CTAAGGCATT	GCCGTCATCT	CGGGAAUAC	ACUTTTTCAG	420
GCTTCTTGC	CTCTGTGTCT	TGGCTGTCT	CTTGGCTGCC	AATCCCATC		469

(2) INFORMATION FOR SEQ ID NO:30:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 359
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: double
 (D) TOPOLOGY: linear
 (ii) MOLECULE TYPE: genomic DNA
 (iii) HYPOTHETICAL: no
 (iv) ANTI-SENSE: no
 (vi) ORIGINAL SOURCE:
 (A) ORGANISM: human
 (x) FEATURE: exon 15 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:

CGGTAGGAAA	GTGATTCTTG	TGTCGACTC	TAGGGCACCC	ACAGCCTCAG	TATGATTGTC	60
CTAGAGGAG	GATGCTCTCT	AAGCCTGCGA	TCTCTTGCTT	CAGACACTG	TTCTTTTCTT	120
GCAGATTCCA	CAGCTGACCA	AACAGGTGTC	CATGGCCAGA	CAAGGCCAGG	TAGATTTCCA	180
ACGAGAGAAA	AAAGAGCTGG	AGGATTGCTT	GGAGCGCATC	ACTGACACAG	GCCAGCGGAA	240
GCTGATTGGG	ACGAGAGACA	CTCGGCAAAAT	GAGGAGGGGG	GCTGTTGAGT	TGTTGGCGGG	300
GGCTTTGTGG	CCTTCTGCTC	CATGGGCAGT	TCTGTGGGTC	GGTGGGCATC	ACACAGCAG	359

(2) INFORMATION FOR SEQ ID NO:31:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 209
 (B) TYPE: nucleic acid
 (C) STRANDEDNESS: double
 (D) TOPOLOGY: linear
 (ii) MOLECULE TYPE: genomic DNA
 (iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 16 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:

GTTGATCGCT	TGGGACGTTT	TTACATTTTT	ATATTCTTTG	TCACTGTCAC	CCAGATCAGA	60
GTCCCTCTGT	TTTTCTTCTC	TTTCAGACTC	AAGAACAGCT	GGAGTTCTTA	GAGAGCTTGA	120
AGCAGGAAC	TCCACCAAGC	CACCGGGAGC	TTGAGGTTCT	GCAAGGCAGC	CTGGAACTT	180
CTGCCAGGT	AAATACCTCC	TTTTTTTTTT				209

(2) INFORMATION FOR SEQ ID NO:32:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 485

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 17 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:

CCCCACATGC	AATCAGTGTG	TCCCCGGGAG	GGATCAGAG	TGGCAGGTTA	AAGACCCATC	60
ACCTTCCCAAG	TCCTTGCAAC	CCGCTGGTGC	CTTGCACCTC	TCXGANGTAG	GGACTGTTTA	120
ACTCAACCAAG	CTCTCCCTTC	TTTCTTTGTG	GTCACTTTTG	CAGTCAGAAAG	CABACTTCCC	180
AGCCCACTTC	GCCGAGCTAG	AGAAGGAGCC	GCACAGCCTC	CTCAGTGGCC	CAGCTCATAG	240
GGAGGAGGAA	TTATCTGCTC	TTCCGGAAGA	ATGCGAGGAC	ATCAGCTCA	AACCTGCCAC	300
CACACAGGCT	CACGGACATG	GACACGAGCC	ACCACCTCTC	AATTCCTACC	GAGGGCCTCT	360
CGCATGCAC	GGAGGCTGGG	ACCAACCCGG	GGCTGCTGAG	AAGGGGTTTG	GGCCCTTGGC	420
CTGATTGTC	AGACATCTCT	TAGGTGTAAT	GCCAGCAGGC	CTTGCATTGC	CTGCACACTC	480
CATGA						485

(2) INFORMATION FOR SEQ ID NO:33:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 468

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 18 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:

TTACTGGCTT	GGACCTCATT	GGCCATGACT	TGAGCTAAGA	TGCTAGAGGC	CCCAGCCAGG	60
TCATCCTGCT	CAGGTCATTT	ATGGAGTCTA	GGCAGACTC	TCACCTCCCT	GGACCATTTT	120

TAGAATCTAT	GTGCCAGCTT	GCCAAAGACC	AACGAAAAT	GCTTCTGGTG	GGGTCCAGGA	180
AGGCTCCCGA	GCAGGTGATA	CAGAGGCCCC	TGAACCAAGT	TGAAGAACCT	CCCTCCATCA	240
GCTGCGCTGG	GTCTGCAGCT	ACACTTGCAG	TTGCCGACCT	GGCAGGGGCC	AGGTCTTAC	300
AGCCTGAGAC	TCTGTTGATG	TTGAATCTCA	TGTGAGACTT	AGCTCAGGGG	CTCTCAGCCT	360
AGCAGCACTT	CAGCATTACC	TTAGGGGGCC	CCAGGGCCCC	TCTTAGATCA	GTTACATGTG	420
GAACTCTGT	GCATTAGTGC	CTATACACTA	GTATTTTAGT	ATTTCTTT		468

(2) INFORMATION FOR SEQ ID NO:34:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 393

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 19 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34:

CACTAGTAG	CTCCTCAT	CAGTCTTA	TTACGAGGA	TGAAGCCAG	TATCAGACT	60
TGCTCTGAC	TTGCTCTG	TTGCTCTCA	CACATCAGCT	CTCTCTCAG	GTCACATCA	120
TTTCCAGCTG	CATCCACAA	CTGGAGAAA	GCTGGAGGCA	GTATCTGCTC	TCTCCAGAG	180
GTAGGATGG	CCAGGACAG	TCTCTCTGG	CTACTCTGG	CCAGACAGGG	TTGAGAGCA	240
CCTGATGCG	GGGATAGTA	CAGTCTCTC	TGCATCAGA	AAGGCACTA	CCCACTCAT	300
ACAGGAAAG	CATCTAGGA	ACTCATAAA	CGGAGGACA	CCCTATGAA	GTGTCACCAT	360
CAACCAGACC	TGAGAACTT	CTCTTCCAA	TCT			393

(2) INFORMATION FOR SEQ ID NO:35:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 421

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 20 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:

GGCTGCTCA	GAGGTAGA	ATGGCCAGG	ACAGTCTCTG	TGGCTTAGTG	ATGCTCCAGC	60
AGGCTTCAG	AGCCTTCAA	TGCGGGGATA	GTGACAGGTC	CTCTCTCATC	AAGAAGGUA	120
TGTAGCCAA	TCATCAGAG	AAGGCACTA	CCCACTCAT	AAAACGGUAG	GAGAGGCTAT	180
CAAGTGTCA	CCATCAACCA	GACCTGACAA	ACTTCTCTTT	CCAACTCTGC	CACACATCAG	240
TGGACTTCTC	CATTCATTA	CCCTGCTGGC	CCACTTGAAC	AGCCACGCCA	TTGCTCATGG	300
TGCCACCACC	TGCTCTAGAG	CCCCACTTGA	CCCTGCGGAC	TGTGAGTACT	GGGECATGAC	360
GGGCTGTTCA	TGGACCAGGG	GACCACTTGG	CTTTTAAAG	TCTCTCTTCC	GCCGGGGCGCA	420
						421

(2) INFORMATION FOR SEQ ID NO:36:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 498

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 21 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36:

AGGCCGAGGC	AGGAGAATCG	CTTGAACTCA	GGAGCCGAG	TTTGCAGTGA	GCCGAGATGG	60
CGCCACTGCA	CTCCAGCCTG	GGCAACAGGA	GCGAGACTTC	ATUTCAAAAA	AAAAGCTGCT	120
ATTGCTTTGT	ATCTTCAGCA	CTGACCGAGG	CCCTTAAGCA	GTATGGCAGG	GAAACCCCTG	180
CTTACCTGGC	CTCCCTCCAG	GAGAGGGGAA	GCCTTGGGAA	TGCUUACAGU	ACAGCCATCA	240
GGAACTGCTT	GAGTAGATC	AAGGCCATCC	CCGCTTACT	TGGAGTAGTA	TCATTGAGGA	300
GCATTGTAT	TCTTCTGCT	GTGCTGCTG	GTGAATGGCC	AGUGAATGAG	TGATGTTCT	360
AGCTAGTTCT	TTCTGCACCT	AGACTTGAT	TCTACAAAGA	CATTGTTAAA	ATTGAAAAAT	420
CTGCCCGGGT	GCAGTGATTT	ATCCCTGTAA	TCCCAGCACT	TTGGGAGGCC	GAGTCAGGAG	480
GATCACTTGA	GGCTAGAC					498

(2) INFORMATION FOR SEQ ID NO:37:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 427

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 22 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 37:

CCCTGTGGCT	TGCAGAAGGT	GTATTGCTGG	TGCTCTCTCTG	CCTTGCCATC	TTGTAAGGCT	60
TACAGATGGC	AGAGGAGAAG	AGACAGGAGG	CCCCAAGGTC	AGTTCAAGCT	TTGTGATGTG	120
TTCAACAGGAG	CTCCTGCCCA	GGGAUTGGA	CATCAAGCAC	GAGGAGCTGG	GGGACCTGGT	180
GGACAAGGAG	ATGGCGGCCA	CTTCAGCTCC	TATTGAAACT	GCCACGGCCA	GAATACAGGT	240
AGGAGGTTCC	TGCAGGATCT	CCTGAAACGA	TGCCTTTGUA	GCATGCCCTC	TGCACCACTG	300
CTCATTAAGC	ATGTCACAGT	CGTTCAATTAA	GGCAATGGCA	ACCCCTTAAG	ACAGAAAUCA	360
GAATTTGCCA	GGCAGAGTGG	CTCATGCTTC	TAACCCCAGC	ACCTTGGGAG	GATCACTTGA	420
CTCCAGG						427

(2) INFORMATION FOR SEQ ID NO:38:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 367

(B) TYPE: nucleic acid
 (C) STRANDEDNESS: double
 (D) TOPOLOGY: linear
 (ii) MOLECULE TYPE: genomic DNA
 (iii) HYPOTHETICAL: no
 (iv) ANTI-SENSE: no
 (vi) ORIGINAL SOURCE:
 (A) ORGANISM: human

(x) FEATURE: exon 23 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 38:

CCCCCTGAAT	ACUUTAGAGT	CTGGATTCCT	TTCTGACTCT	CTCAAGAGTG	TGGGCAAGGA	60
CTTGGXXACT	TCCAGATTCA	GGTTTCCCAG	CTACCACACC	ATGTTGGACT	GAAAGTATAG	120
TAAACATTA	GTGGATCCTT	AATATTCAAG	GCACATTTAG	AAACCATGCT	TCTTTTTCAC	180
AGGAGATGCT	CAGCAATTC	CGAGCAGGAG	ACACACCACT	CAAAATTGGAG	GTGANTGAAA	240
GGTCGGTCTG	AGCGGCATTC	TGGGACCTAG	GGGAGCAGGA	TCGTCTTTTC	TGACATTTGT	300
CTATACTTTC	CATACTTATT	AGGGAATTAG	AGGAGAGCAG	TACCAGCCAC	CCGCAACXXC	360
TGAGTTG						367

(2) INFORMATION FOR SEQ ID NO:39:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 502

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 24 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:

CCCCGCGGAA	TCTTCCAGCA	ACCTCAGCAC	CTTCTTACC	TCCCTTTCCC	ATTCCAAGCT	60
TGCCCTTTGGC	TAGGAGTGGG	GAAGACAACC	GTCCGTGTCA	TTGATCTTGG	AGCTTGATCT	120
CAGTGATACC	TCCACTTCTT	TGTSTGGCAG	GATCCCTGCT	TCCGTATCCA	GCCTCATGCA	180
AGCTATTGAC	GTGTCATCG	TGGCCTCTAA	GGACCTCCAG	AGAGAGATTG	TGGAGAGCGC	240
CAGCGTGAGC	GTGGGTGTGG	GGCCTGGGCA	GGAAGAGGAG	GCACTGGTGA	CACACTCCCG	300
CTCCAACGGA	CTCTGTGATG	CTGCGGTCTT	ACTCTGTGTG	TCCACCTGAG	TACAGAGCAG	360
CCACTCCTCT	AGATATCAGC	AGAGGCCCTC	GGCAGAGGTC	AGAGCTCCAG	GACCTCCCCA	420
GAGGGTGGCC	AGGCACTGCT	CCCACTCCCA	GCTCCCTTGG	CACAGGCACA	CATTGTTGGA	480
ACTTGCCTGG	GGAGCCCTTT	TT				502

(2) INFORMATION FOR SEQ ID NO:40:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 437

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 25 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 40:

TTTTGGTCTC	TGAATCTTCT	TCTTTTTTGT	AAATGGGAA	TACTAATGCT	TATGTCTCAG	60
AGTTACTATG	AGGATGATTT	CCGATAATAT	ATCTATAAAA	CCACCTGCCA	TATAGTACAT	120
CCTCAATNAA	AGGTGGCTAT	TACTATTTTT	TATTTCCCTA	GGGTACAGCA	TCCCTTAAAG	180
AGTTTATAGC	CAAGAACTCT	CCATGGACAG	AACTACTTAT	CTCAGCCTCC	AAGGCTGTGG	240
GCTGGGGAGC	CATGTCTATG	GTGTAAGTAT	CTATTGGTAC	CAGGGGTCTT	CCCATGACCC	300
CTCTTCCATT	GATCCACTCC	AAACAATAGC	TAACGAGGCA	AAAAAAATC	TGTCCTTTAG	360
AAATAAATA	TTGATCAGGA	AGTCAATAGG	ACCGAGTTTA	CAGGGGAGCC	TGGCTCTCCC	420
AGGGGACACA	GAGCAGG					437

(2) INFORMATION FOR SEQ ID NO:41:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 351

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 26 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:

GGGAGCCTGG	CTCTCCGAGG	GGACACAGGG	CAGGCACTCT	CCCCCTCCTG	TTTAGCCAGG	60
GGCGATGGGG	TGGTCGGAG	GTGCAATGT	GGAGGAGTTG	CAGCTCATTT	CCCCCTAACC	120
TAGTCCCTCT	TCTCGTTTTC	CATCAGGGAT	GCAGCTGATC	TGCTGGTACA	AGGCAGAGGG	180
AAATTTGAGG	AGCTAATGCT	GTGTTCTCAT	CAATTTGCTG	CTAGCCACAG	CCAGCTTGTG	240
GCTGCATCCA	AGGTAGGACC	TGGCTGGACC	TCCTAGGACG	CTGGAGAGCC	TGCTTAGAGA	300
GTACTAGGCT	ACCTTAAGA	GTACTTGGCT	GAGTAAAGCA	CTACTTGGCT	G	351

(2) INFORMATION FOR SEQ ID NO:42:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 418

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 27 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:

CTTTTTATAT	GATAGATATG	TGAGGAGCTG	ACTATACTCA	GCAGATTTTG	AGAAGCTGAT	60
TGGTGATTGC	CGTGTGGCCC	ACATATCTTT	GCTAAGAACC	ATCAUAGCAA	TTATCTGATT	120
CAGTCTTGT	TGCTCTAGGT	GTGTATGAA	CCATAATCTG	CTTTGTCTCTG	GTAGGTGAAA	180

GCTGATAAGG	ACAGCCCCAA	CCTAGCCCAG	CTGCAGUAGG	CCTCTCGGGG	AGTGAACCAg	240
GCCACTGCCG	GCGTTGTGGC	CTCAACCAT	TCCGECANAT	CACAGATCNA	MCAGACAGGT	300
AGCCTTTCCA	AMGGGACCCT	TTTCTTACCC	ACCTGTITGA	GCTCTTCTCT	GCATCCTTCC	360
CTGTGATCCC	AACUAAATCC	CACAGGACTG	TGTCTAAAT	CTTTCATAT	TTTCATCT	418

(2) INFORMATION FOR SEQ ID NO:43:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 279

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 28 of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:

TTTCCACAGA	GCATTGCCAT	TGGCTGCTTC	TCAGGTGCCA	GTCACCCACG	CTAGATTTTG	60
ATGAGACCTT	CTTGTTTCCA	TCCCTTGCACA	CAACATGGAC	TTCTCAGCA	TGACGUTGAC	120
ACAGATCAAA	CGCCAAAGACA	TGGATTCTCA	GGTTAGGCTG	CTAGAGCTAC	AAANTGATTT	180
GCAGAAGGAC	CTCAAAAAC	TGGGACAGCT	TGGGAAAAG	CACTACGAGC	TTGCTGCTGT	240
TGCTGAGGGC	TGGGAAGAAC	GTAAGCTGAC	TCAAAGGAT			279

(2) INFORMATION FOR SEQ ID NO:44:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 3715

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: genomic DNA

(iii) HYPOTHETICAL: no

(iv) ANTI-SENSE: no

(vi) ORIGINAL SOURCE:

(A) ORGANISM: human

(x) FEATURE: exon 29 and partial cds of HIP1

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 44:

ABCATAAAAT	ATCATTGTCT	TTTAGGACCA	GAGGCATCTC	CACCTACACT	GCAAGAAGTG	60
GTACCCGAAA	AAGATAGAG	CCAAACCAAC	ACCCCATATG	TCAGTGTAAA	TCTTGTATAC	120
CTATCTCGTG	TGTATTATTT	CCCCAGCCAC	AGGCCAAATC	CTTGGAGTCC	CAGGGGCAGC	180
CACACCACTG	CCATTACCCA	GTGCCGAGGA	CATGCATGAC	ACTTCCCAA	GACTCCCTCC	240
ATAGCGACAC	CCTTTCTCTT	TGGACCCATG	GTGATCTCTG	TTCTTTTCCC	GCCTCCCTAG	300
TTAGCATCCA	GGCTGGCTAG	TGCTGCCCAT	CAGCANGCCT	AGGTACGAAG	AGGAGTGTCT	360
GGGGGCGAGG	CCACTCAACA	CAGAGGACCA	ACATCCAGTC	CTGCTGAUTA	TTTGACCCCC	420
ACAACATCG	GTATCCTTAA	TAGAGGAGCT	GCTTCTTCTT	TCTTGACAGC	TTGGAAGAGG	480
AAGATCTTAT	GCCTTTTCTT	TTCTGTTTTC	TTCTCASTCT	TTTCAGTTTC	ATCATTTCCT	540
CAAACCTGTG	AGCATCAGAG	CCCTGATGGA	TTCCAAACCA	GGACACTACC	CTGAGATCTG	600
CACAGTCAGA	AGGACGGCAG	BAGTGTCTTG	GCTCTCAATC	CCAAAGCCAT	TCTCCCTCTC	660
TTTGGGCGAT	CCCATGGATT	TCCACTGCTT	CTTATGGTGG	TTGGTTGGGT	TTTGTGTTT	720
TGTCTTTTCT	TTTAAAGTTT	CATCACATA	GCCAACTCTC	CCAAAGCCCA	CACCCCTGGG	780
GCTGAGTCTC	CAGGGCCCCC	CAACTGTGGT	AGCTCCAGCG	ATGGTGCTGC	CCACCTCTCT	840

CGGTGCTCCA	TCTCCGCTTC	CACACTGACC	AAGTGCTGGC	CCACCCAGTC	CATGCTCCAG	900
GGTCAGGCGG	AGCTGCTGAG	TGACACCTTT	CCTCAAAAAG	CAGAAGGAGA	GTGAGTGCCT	960
TTCCCTCCTA	AAGCTGAATC	CCGGCCGAAA	GCCTCTGTCC	GCCTTTACAA	GGGAGAAGAC	1020
AACAGAAAGA	GGGACAGAGG	GGTTCACADA	GGCCAGTTCC	CGTGACGAGG	CTCAAAAACCT	1080
TGATCACAATG	CTTCAATGGA	GCTGGTGAGA	TCAACAACAC	TACTTCCCTG	CCGGAATGAA	1140
CTGTCCGTGA	ATGGTCTCTG	TCAAGCGGGC	CGTCTCCCTT	GGCCACGAGA	CGGAGTGTGG	1200
GAGTGATTCC	CAACTCCCTT	CTGTAGACCT	CTGCCCTGGC	ATCCTCTTGA	ATAGGAAGAT	1260
CGTTCCACTT	TCTACGCAAT	TGACAAACCC	GGAGAGTCAG	ATGCAATTTG	TCCCATCAGG	1320
GAAGAACCCCT	ATACTTGCTT	TCTTACCCCT	AGTATTTTAT	ACTAACCTCC	CTTAAACAGC	1380
AACACCCCTAC	AAGGAGATGC	TTCGAGCAAT	CAGAACTTCA	GTGTGACTTC	TAGCAAAAGCT	1440
CATCTTTCTG	CCCGGCTACA	TCAACCTTCA	AGATTCAGAA	GAAAGCCAAAG	GTGCTGGACT	1500
GTTACTGACT	TGGATCCCAA	AGCAAGGAGA	TCATTTGGAG	CTCTTGGGTC	AGAGAAATG	1560
AGAAAGGACA	GAGCCAGCGG	CTCCAACTCC	TTTCAGCCAC	ATGCCCCAGC	CTCTCCTCTG	1620
CCTGTGGACA	GGAAGAGGAC	AGAGGGCACA	TGACAGCCTT	GCCAGGGATG	GGCAGCCCAA	1680
CAGCACTTTT	CCTCTCTTAG	ATGGACCCCA	GCATTTAAGT	GACCTTCTGA	TCTTGGGAAA	1740
ACAGCGTCTT	CTTCTTTTAT	CTATACCAAAC	TCATTTGGTG	TAGCCATCAA	GTACTTCCCA	1800
GGATCTGCTC	CAACAGAAATA	TTGCTAGGTT	TTGCTACATG	ACGGGTGTGT	AGAATTCTGT	1860
TTGATCACTG	TGAACCAACU	CCCAATCTCC	TAGCCCAAGC	CCCTCCCCAA	CTCCCTCTCT	1920
GTGCATTTTC	TAAGTGGGAC	ATTCAAAAAA	CTCTCTCCCA	GGACCTCGGA	TGACCAATAT	1980
CAGACGTGTG	ACCTCCATAC	TGGGTAAAGC	AAGTATTCAG	ACTAGAAATT	GGGAGTCTTT	2040
AATGTTGAAT	GCCTCTTTCT	GCTTAGTATT	TTTTTGATTC	AAGGCTCAGA	AGGAATGGTG	2100
CGTGGCTTCC	CTGTCCCACT	TGTGGCAACT	AAAGCAATCC	CTGTCTTTCT	GATGCCGGCT	2160
AACATTTCCA	AAAGTGGCTA	CTCTTCACTT	CTAGATCTCA	GCCATTTCTA	CTCATATGTT	2220
CCCAATTAAC	AAAGGGTGGC	CGGGCACAGT	GGCTCAGGCC	TGTAATCCCA	CCACTTTGAC	2280
AGGCTGAGGT	GGTAGGATCA	CTTGAGGTCA	CCAGTTCAAG	ACCAGCCTGT	CCAACATGCT	2340
GAAACCCCAA	TCTCTACTAA	AAATACCAAA	AATTAGCCGA	GGTAGTGAAC	GGTCCCTCTT	2400
AAATCCACCT	ACTCAGGAGG	CTGAGACAGG	ACCAATACCT	GAACCCCAAG	GGCAGAGGTT	2460
CCAGTGAGCT	GAGATCACCC	CATTGTACTC	CAGCCTGGGC	AACAAGAGCA	AAACTCCCTC	2520
TCAAAAAAAA	AAAAAATTA	CAATGGGGG	AAACAGTCTA	CTGTATTTGA	TCAATTTAAG	2580
ATTCTCTGCC	CAGCUGGGCA	CAGTGGGGCA	TGCCCTGTAT	CCCAGAAATT	TGGGAGGCCA	2640
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AAATTTATCT	GAATGATCCT	CTCTCTAAAA	AGAGGCCACA	GAATGTGTTA	AAAACCTTCA	3060
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GTGTCACTAC	AGGAAGTAGC	TGTCAGATGT	CACATTAATTA	CTTTCTGTAT	AGCTCAGATT	3180
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GACTAGCCTG	GCCAAACAAG	TGAAACCCCG	TCCTCTACTA	AAATACAAAA	ATTAGCCCGG	3660
CATGGTGCCA	GGCCCTATAA	ATCCCAAGTA	CTGGGAGGCG	TGAGCCAGCA	GAATC	3715

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(21) International Application Number: PCT/US99/11743 (22) International Filing Date: 27 May 1999 (27.05.99) (30) Priority Data: 09/085,199 27 May 1998 (27.05.98) US (71) Applicants (for all designated States except US): UNIVERSITY OF BRITISH COLUMBIA [CA/CA]; University Industry Liaison Office, IRC Building - Room 331, 2194 Health Sciences Mall, Vancouver, British Columbia V6T 1Z3 (CA). MERCK FROSST CANADA & CO. [CA/CA]; PO/CP 1005, Pointe Claire-Dorval, Québec H9R 4P8 (CA). (72) Inventors; and (75) Inventors/Applicants (for US only): KALCHMAN, Michael [CA/CA]; #1403-900 Yonge Street, Toronto, Ontario M4W 3P5 (CA). HAYDEN, Michael, R. [US/CA]; 4484 West Seventh, Vancouver, British Columbia V6R 1W9 (CA). HACKAM, Abigail [CA/CA]; 1420 West 11th Avenue, Vancouver, British Columbia V6H 1L2 (CA). CHOPRA, Vikramjit [CA/CA]; Suite 210, 2475 Blenheim Street, Vancouver, British Columbia V6K 4N7 (CA). NICHOLSON, Donald, W. [CA/CA]; 18-750 Milton Street, Montréal, Québec H2X 1W4 (CA). VALLAINCOURT, John, P. [CA/CA]; 18022 Amalfi Street, Québec, Québec H9K 1N7		(CA). RASPER, Dita, M. [CA/CA]; Apartment #7, 16203 Pierrefonds Boulevard, Pierrefonds, Québec H9H 4S8 (CA). (74) Agent: COPPOLA, Joseph, A.; Merck & Co., Inc., P.O. Box 2000, Rahway, NJ 07065-0907 (US). (81) Designated States: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG). Published <i>With international search report.</i> (88) Date of publication of the international search report: 20 April 2000 (20.04.00)

(54) Title: APOPTOSIS MODULATORS THAT INTERACT WITH THE HUNTINGTON'S DISEASE GENE**(57) Abstract**

A family of proteins, including a specific human protein designated as HIP1, has been identified that interact differently with the gene product of a normal (16 CAG repeat) and an expanded (>44 CAG repeat) HD gene. Expression of the HIP1 protein was found to be enriched in the brain. Analysis of the sequence of the HIP1 protein indicated that it includes a death effector domain (DED), suggesting an apoptotic function. Thus, it appears that a normal function of Huntingtin may be to bind HIP1 and related apoptosis modulators, reducing its effectiveness in stimulating cell death. Since expanded huntingtin performs this function less well, there is an increase in HIP1-modulated cell death in individuals with an expanded repeat in the HD gene. This understanding of the likely role of huntingtin and HIP1 or related proteins (collectively "HIP-apoptosis modulating proteins") in the pathology of Huntington's disease offers several possibilities for therapy. First, because the function of huntingtin apparently depends at least in part on the ability to interact with HIP-apoptosis modulating proteins, added expression (e.g., via gene therapy) of normal (non-expanded) huntingtin or of the HIP-binding region of huntingtin should provide a therapeutic benefit. Other DED-interacting peptides could also be used to mask and reduce the interaction of HIP-apoptosis modulating proteins with the death signaling complex. Alternatively, a mutant form of HIP-protein from which the DED has been deleted might be introduced, for example using gene therapy techniques. Because HIP-apoptosis modulating proteins have been shown to self-associate, a protein with a deleted DED may compete with endogenous HIP-protein in the formation of these associations, thereby reducing the amount of apoptotically-active HIP-protein.

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According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

U.S. : 536/23.5; 435/6; 530/350

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

USPAT, CAPLUS, MEDLINE

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WANKER et al., HIP-I: A Huntingtin Interacting Protein Isolated by the Yeast Two-hybrid System. Human Molecular Genetics. March 1997, Vol. 6, No. 3, pages 487-495, see entire document.	1-15

☐ Further documents are listed in the continuation of Box C.

☐ See patent family annex.

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